ANSWERS TO PROBLEMS

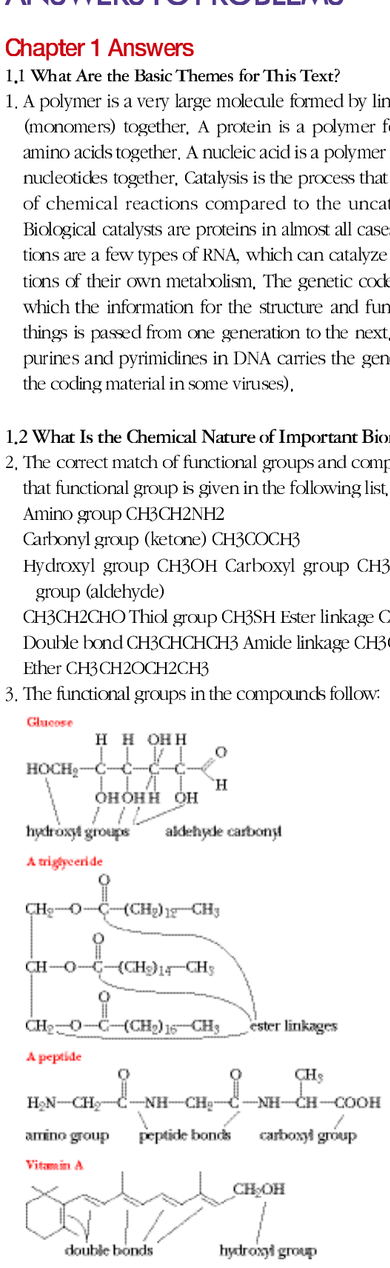
Chapter 1 Answers

1.1 What Are the Basic Themes for This Text?

1. A polymer is a very large molecule formed by linking smaller units (monomers) together. A protein is a polymer formed by linking amino acids together. A nucleic acid is a polymer formed by linking nucleotides together. Catalysis is the process that increases the rate of chemical reactions compared to the uncatalyzed reaction. Biological catalysts are proteins in almost all cases; the only exceptions are a few types of RNA, which can catalyze some of the reactions of their own metabolism. The genetic code is the means by which the information for the structure and function of all living things is passed from one generation to the next. The sequence of purines and pyrimidines in DNA carries the genetic code (RNA is the coding material in some viruses).

1.2 What Is the Chemical Nature of Important Biomolecules?

2. The correct match of functional groups and compounds containing that functional group is given in the following list.
   Amino group $CH_3CH_2NH_2$
   Carbonyl group (ketone) CH_3COCH_3
   Hydroxyl group CH_3OH Carboxyl group CH_3COOH Carbonyl group (aldehyde)
   CH_3CH_2CHO Thiol group CH_3SH Ester linkage $CH_3COOCH_2CH_3$
   Double bond $CH_3CHCHCH_3$ Amide linkage $CH_3CON(CH_3)2$
   Ether $CH_3CH_2OCH_2CH_3$
3. The functional groups in the compounds follow:



4. Before 1828, the concept of vitalism held that organic compounds could only be made by living systems and were beyond the realm of laboratory investigations. Wohler's synthesis showed that organic compounds, like inorganic ones, did not require a vitalistic explanation, but rather they obeyed the laws of chemistry and physics and thus were subject to laboratory investigation. Subsequently, the concept was extended to the much more complex, but still testable, discipline of biochemistry.
5. Urea, like all organic compounds, has the same molecular structure whether it is produced by a living organism or not.
6. 

7. Five; 7 if the two cyclopropane derivatives are allowed.
8. Thirteen different alcohols, 11 aldehydes/ketones, and 10 each epoxides and ethers.

1.3 What Can Biochemistry Say About Possible Origins of Life?

9. It is generally believed that carbon is the likely basis for all life forms, earth-bound or extraterrestrial.
10. Eighteen residues would give 20^{18} or $2{,}6 \times 10^{23}$ possibilities. Thus 19 residues would be necessary to have at least Avogadro' s number ($6{,}022 \times 10^{23}$) of possibilities.
11. The number is 4^{40} or $1{,}2 \times 1024$, which is twice Avogadro's number.
12. RNA is capable of both coding and catalysis.
13. Catalysis allows living organisms to carry out chemical reactions much more efficiently than without catalysts.
14. Two of the most obvious advantages are speed and specificity; they also work at constant temperature or produce little heat.
15. Coding allows for reproduction of cells.
16. With respect to coding, RNA-type polynucleotides have been produced from monomers in the absence of either a preexisting RNA to be copied or an enzyme to catalyze the process. The observation that some existing RNA molecules can catalyze their own processing suggests a role for RNA in catalysis. With this dual role, RNA may have been the original informational macromolecule in the origin of life.
17. It is unlikely that cells could have arisen as bare cytoplasm without a plasma membrane. The presence of the membrane protects cellular components from the environment and prevents them from diffusing away from each other. The molecules within a cell can react more easily if they are closer to each other.

1.4 How Do Prokaryotes and Eukaryotes Differ in Levels of Organization?

18. Five differences between prokaryotes and eukaryotes are: (1) Prokaryotes do not have a well-defined nucleus, but eukaryotes have a nucleus marked off from the rest of the cell by a double membrane. (2) Prokaryotes have only a plasma (cell) membrane; eukaryotes have an extensive internal membrane system. (3) Eukaryotic cells contain membrane-bounded organelles, while prokaryotic cells do not. (4) Eukaryotic cells are normally larger than those of prokaryotes. (5) Prokaryotes are single-celled organisms, while eukaryotes can be multicellular as well as single-celled.
19. Protein synthesis takes place on ribosomes in both prokaryotes and eukaryotes. In eukaryotes, ribosomes may be bound to the endoplasmic reticulum or found free in the cytoplasm; in prokaryotes, ribosomes are only found free in the cytoplasm.

1.5 What Are the Main Structural Features of Prokaryotic Cells?

20. It is unlikely that mitochondria would be found in bacteria. These eukaryotic organelles are enclosed by a double membrane, and bacteria do not have an internal membrane system. The mitochondria found in eukaryotic cells are about the same size as most bacteria.

1.6 What Are the Main Structural Features of Eukaryotic Cells?

21. See Section 1.6 for the functions of the parts of an animal cell, which are shown in Figure 1.10(a).
22. See Section 1.6 for the functions of the parts of a plant cell, which are shown in Figure 1.10(b).
23. In green plants photosynthesis takes place in the membrane system of chloroplasts, which are large membrane-enclosed organelles. In photosynthetic bacteria, there are extensions of the plasma membrane into the interior of the cell called chromatophores, which are the sites of photosynthesis.
24. Nuclei, mitochondria, and chloroplasts are bounded by a double membrane.
25. Nuclei, mitochondria, and chloroplasts all contain DNA. The DNA found in mitochondria and in chloroplasts differs from that in the nucleus.
26. Mitochondria carry out a high percentage of the oxidation energy-releasing reactions of the cell. They are the primary sites of ATP synthesis.
27. The Golgi apparatus is involved in binding carbohydrates to proteins and in exporting substances from the cell. Lysosomes contain hydrolytic enzymes, peroxisomes contain catalase (needed for the metabolism of peroxides), and glyoxysomes contain enzymes needed by plants for the glyoxylate cycle. All these organelles have the appearance of flattened sacs bounded by a single membrane.

1.7 How Do We Classify Organisms: Five Kingdoms or Three Domains?

28. Monera include bacteria (e. g., E. coli) and cyanobacteria. Protista include Euglena, Volvox, Amoeba, and Paramecium. Fungi include molds and mushrooms. Plantae include clubmosses and oak trees. Animalia include spiders, salmon, rattlesnakes, robins, and dogs.
29. The kingdom Monera consists of prokaryotes. The other four kingdoms consist of eukaryotes.
30. The kingdom Monera is divided into the domains Eubacteria and Archaea on the basis of biochemical differences. The domain Eukarya consists of the four kingdoms of eukaryotic organisms.

1.8 Is There Common Ground for All Cells?

31. The major advantage is that of having compartments (organelles) with specialized functions (and thus division of labor). Another advantage is that cells can be much larger without surface area to volume considerations being critical because of compartmentalization.
32. See the discussion of the endosymbiotic theory in Section 1.8.
33. See Exercise 32. The division of labor in cells gives rise to greater efficiency and a larger number of individuals. This in turn allows more opportunity for evolution and speciation.

1.9 How Do Cells Use Energy?

34. Processes that release energy are favored.

1.10 What Is the Connection between Energy and Change?

35. The term "spontaneous" means energetically favored. It does not necessarily mean fast.

1.11 What Is the Criterion for Spontaneity in Biochemical Reactions?

36. The system is the nonpolar solute and water, which become more disordered when a solution is formed; ΔSsys is positive but comparatively small. The ΔSsurr is negative and comparatively large because it is a reflection of the unfavorable enthalpy change for forming the solution (ΔHsys).
37. Processes (a) and (b) are spontaneous, whereas processes (c) and (d) are not. The spontaneous processes represent an increase in disorder (increase in the entropy of the universe) and have a negative $\Delta G°$ at constant temperature and pressure, while the opposite is true of the nonspontaneous processes.
38. In all cases, there is an increase in entropy, and the final state has more possible random arrangements than the initial state.
39. Since the equation involves multiplication of ΔS by T, the value of ΔG is temperaturedependent.
40. If one considers entropy to be a measure of dispersion of energy, then at higher temperatures, it is logical that molecules would have more possible arrangements due to increased molecular motion.
41. Assuming the value of DS is positive, an increase in temperature will increase the - contribution of the entropy component to the overall energy change.
42. The heat exchange, getting colder, reflects only the enthalpy or ΔH component of the total energy change. The entropy change must be high enough to offset the enthalpy component and add up to an overall $-\Delta G$.
43. Entropy would increase. There are more ways that two molecules, ADP and Pi, can be randomized than a single molecule, ATP.

1.12 What Is the Connection between Thermodynamics and Life?

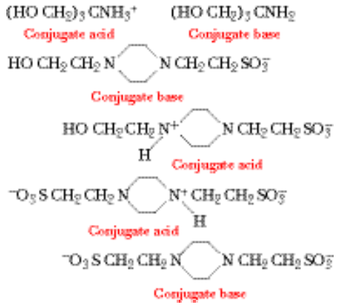
44. The lowering of entropy needed to give rise to organelles leads to higher entropy in the surroundings, thus increasing the entropy of the universe as a whole
45. Compartmentalization in organelles brings components of reactions into proximity with one another. The energy change of the reaction is not affected, but the availability of components allows in to proceed more readily.
46. DNA would have higher entropy with the strands separated. There are two single strands instead of one double strand, and the single strand have more conformational mobility.
47. See the answer to exercise 43. It is still unlikely that cells could have arisen as bare cytoplasm, but the question of proximity of reactants is more to the point here than the energy change of a given reaction.
48. It would be unlikely that cells of the kind we know would have evolved on a gas giant. The lack of solids and liquids on which aggregates could form would make a large difference.
49. The available materials differ from those that would have been found on earth, and conditions of temperature and pressure are very different.
50. Mars, because of conditions more like those on earth.
51. A number of energetically favorable interactions drive the process of protein folding, ultimately increasing the entropy of the universe.
52. Photosynthesis is endergonic, requiring light energy from the sun. The complete aerobic oxidation of glucose is exergonic and is a source of energy for many organisms, including humans. It would be reasonable to expect the two processes to take place differently in order to provide energy for the endergonic one.

Chapter 2 Answers

2.2 What Is a Hydrogen Bond?

1. Proteins and nucleic acids have hydrogen bonds as an important part of their structures.
2. Replication of DNA and its transcription to RNA requires hydrogen bonding of complementary bases to the DNA template strand.
3. The C—H bond is not sufficiently polar for greatly unequal distribution of electrons at its two ends. Also, there are no unshared pairs of electrons to serve as hydrogen bond acceptors.
4. Many molecules can form hydrogen bonds. Examples might be H_2O, CH_3OH, NH_3.
5. For a bond to be called a hydrogen bond, it must have a hydrogen covalently bonded to O, N, or F. This hydrogen then forms a hydrogen bond with another O, N, or F.
6. In a hydrogen-bonded dimer of acetic acid the —OH portion of the carboxyl group on molecule 1 is hydrogen-bonded to the—C=O portion of the carboxyl group on molecule 2, and vice versa.
7. Glucose = 17 and sorbitol =18, ribitol + 15; each alcohol group can bond to 3 waters and the ring oxygen binds to two. The sugar alcohols bind more than the corresponding sugars.)
8. Hypoventilation decreases the pH of blood. (See BC on p. 54)
9. Aspirin is electrically neutral at the pH of the stomach and can pass the membrane more easily there than in the small intestine.
10. Positively charged ions will bind to nucleic acids as a result of electrostatic attraction to the negatively charged phosphate groups.
11. The unique fitness of water for forming hydrogen bonds determines the properties of many important biomolecules. Water can also act as an acid and a base, giving it great versatility in biochemical reactions.
12. If atoms did not differ in electronegativity, there would be no polar bonds. This would drastically affect all reactions that involve functional groups containing oxygen or nitrogen. That is most of biochemistry.

2.3 What Are Acids and Bases?

13. $(CH_3)_3NH^+$ (conjugate acid)
    $(CH3)3N$ (conjugate base)
    $^+H_3N—CH_2—COOH$ (conjugate acid)
    $^+H_3N—CH_2—COO^-$ (conjugate base)
    $^+H_3N—CH_2—COO^-$ (conjugate acid)
    $H_2N—CH_2—COO^-$ (conjugate base)
    $^-OOC—CH_2—COOH$ (conjugate acid)
    $^-OOC—CH_2—COO^-$ (conjugate base)
    $^-OOC—CH_2—COOH$ (conjugate base)
    $HOOC—CH_2—COOH$ (conjugate acid)
14. 

15.
(a) The numerical constant equal to the concentration of the products of the dissociation divided by the concentration of the undissociated acid form: $([H^+][A^-])/[HA]$
(b) The qualitative or quantitative description of how much acid (HA) dissociates to hydrogen ion.
(c) The property of a molecule that has both a polar region and an nonpolar region.
(d) The amount of acid or base that can be added to a buffer before experiencing a sharp pH change.
(e) The point in a titration curve where the added acid or base equals the amount of buffer originally present.
(f) The property of a molecule that is readily soluble in water (i.e. water loving)
(g) The property of a molecule that is insoluble in water (i.e. water hating).
(h) The property of a molecule that is not soluble in water. The property of a covalent bond where there is even sharing of electrons and no dipole moments (partial charges).
(i) The property of a molecule that is soluble in water. The property of a covalent bond where the electrons are not shared evenly and dipole moments (partial charges) exist.
(j) An experiment whereby acid or base is added stepwise to a compound and the pH is measured as a function of the added substance.

2.4 What is pH, and What Does It Have To Do with the Properties of Water?

16. The definition of pH is -log$[H^+]$. Due to the log function, a change in concentration of
10 will lead to a change in pH of 1. The log of 10 is one. The log of 100 is 2, etc.
17. Blood plasma, pH 7.4 $[H^+]$ = 4 0 × l0-8 M
    Orange juice, pH 3.5 $[H^+]$ = 3.2 × 10-4 M
    Human urine, pH 6.2 $[H^+]$ = 6.3 × 10-7 M
    Household ammonia, pH 11.5 $[H^+]$ = 3.2 × 10-l2 M
    Gastric juice, pH 1.8 $[H^+]$ = 1.6 × l0-2 M
18.
    Saliva, pH 6.5 $[H^+]$ = 3.2 × 10-7 M
    Intracellular fluid (liver), pH 6.9 $[H^+]$ = 1.6 × 10-7 M
    Tomato juice, pH 4.3 $[H^+]$ = 5.0 × 10-5 M
    Grapefruit juice, pH 3.2 $[H^+]$ = 6.3 × 10-4 M
19.
    Saliva, pH 6.5 $[OH^-]$ = 3.2 × 10-8 M
    Intracellular fluid (liver),pH 6.9 $[OH^-]$ = 7.9 × 10-8 M
    Tomato juice, pH 4.3 $[OH^-]$ = 2.0 × 10-10 M
    Grapefruit juice, pH 3.2 $[OH^-]$ = 1.6 × 10-11 M

2.6 What Are Buffers, and Why Are They Important?

20. The pK of the buffer should be close to the desired buffer pH, and the substance chosen should not interfere with the reaction being studied.
21. The useful pH range of a buffer is one pH unit above and below its p*K*a.
22. Use the Henderson-Hasselbalch equation
    $pH = pK\,a^+ \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $5.00 = 4.76 + \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $0.24 = \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $[CH_3COO^-]$ = inverse log of 0.24 = 1.7
    $[CH_3COOH]$ 1
23. Use the Henderson-Hasselbalch equation
    $pH = pK\,a^+ \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $4.00 = 4.76 + \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $-0.76 = \log ([CH_3COO^-])$
    $([CH_3COOH])$
    $[CH_3COO^-]$ = inverse log of -0.76 = 0.17
    $[CH_3COOH]$ 1
24. Use the Henderson-Hasselbalch equation
    $pH = pK\,a^+ \log ([TRIS])$
    $([TRIS\text{-}H^+])$
    $8.7 = 8.3 + \log ([TRIS])$
    $([TRIS\text{-}H^+])$
    $0.4 = \log ([TRIS])$
    $([TRIS\text{-}H^+])$
    $[TRIS]$ = inverse log of 0.4 = 2.5
    $[TRIS\text{-}H^+]$ 1
25. Use the Henderson-Hasselbalch equation
    $pH = pK\,a^+ \log ([HEPES])$
    $([HEPES\text{-}H^+])$
    $7.9 = 7.55 + \log ([HEPES])$
    $([HEPES\text{-}H^+])$
    $0.35 = \log ([HEPES])$
    $([HEPES\text{-}H^+])$
    $[HEPES]$ = inverse log of 0.35 = 2.2
    $[HEPES\text{-}H^+]$ 1
26. At pH 7.5, the ratio of $[HPO_4^{2-}]/[H2PO_4^-]$ is 2/1 (p*K*a of $H_2PO_4^-$ = 7.2), as calculated using the Henderson-Hasselbalch equation. K_2HPO_4 is a source of the base form, and HC1 must be added to convert one-third of it to the acid form, according to the 2/1 base/acid ratio. Weigh out 8.7 grams of K_2HPO_4 (0.05 moles, based on a formula weight of 174 grams/mole), dissolve in a small quality of distilled water, add 16.7 mL of 1 M HC1 (gives 1/3 of 0.05 moles of hydrogen ion, which converts 1/3 of the 0.05 moles of HPO_4^{2-} to $H_2PO_4^-$) and dilute the resulting mixture to one liter.
27. A 2/1 ratio of the base form to acid form is still needed, because the pH of the buffer is the same in both problems. NaH_2PO_4 is a source of the acid form, and NaOH must be added to convert two thirds of it to the base form. Weigh out 6.0 grams of NaH_2PO_4 (0.05 moles, based on a formula weight of 120 grams/mole), dissolve in a small quantity of distilled water, add 33.3 mL of 1 M NaOH (gives 2/3 of 0.05 moles of hydroxide ion, which converts 2/3 of the 0.05 moles of $H_2PO_4^-$ to HPO_4^{2-}) and dilute the resulting mixture to one liter.
28. After mixing the buffer solution (100 mL) contains 0.75 M lactic acid and 0.25 M sodium lactate. The p*K*a of lactic acid is 3.86. Use the Henderson-Hasselbalch equation
    $pH = pK\,a^+ \log ([CH_3CHOHCOO^-])$
    $([CH3CHOHCOOH])$
    $pH = 3.86 + \log ([CH_3CHOHCOO^-])$
    $([CH_3CHOHCOOH])$
    $pH = 3.86 + \log (0.25\ M/0.75\ M)$

pH = 3,86 + (-0.48)
pH = 3,38

29. After mixing the buffer solution (100 mL) contains 0,25 M lactic acid and 0,75 M sodium lactate. The p*K*a of lactic acid is 3,86. Use the Henderson-Hasselbalch equation
pH = p*K* a$^+$ log ([$CH_3CHOHCOO^-$])
([$CH_3CHOHCOOH$])
pH = 3,86 + log ([$CH_3CHOHCOO^-$])
([$CH_3CHOHCOOH$])
pH = 3,86 + log (0,75 M/0,25 M)
pH = 3,86 + (0,48)
pH = 4,34
30. Use the Henderson-Hasselbalch equation
pH = p*K* a$^+$ log ([CH_3COO^-])
([CH_3COOH])
pH = 4,76 + log ([CH_3COO^-])
([CH_3COOH])
pH = 4,76 + log (0,25 M)
(0,10 M)
pH = 4,76 + 0,40
pH = 5,16
31. Yes, it is correct, calculate the molar amounts of the two forms and insert into the Henderson-Hasselbalch equation. 2,02 g = 0,0167 mol and 5,25 g = 0,0333 mol)
32. The solution is a buffer because it contains equal concentrations of TRIS in the acid and free amine forms. When the two solutions are mixed, the concentrations of the resulting solution (in the absence of reaction) are 0,05 M HCl and 0,1 M TRIS because of dilution. The HCl reacts with half the TRIS present, giving 0,05 M TRIS (protonated form) and 0,05 M TRIS (free amine form).
33. Any buffer that has equal concentrations of the acid and basic forms will have a pH equal to its p*K*a. Therefore the buffer from exercise 32 will have a pH of 8,3.
34. First calculate the moles of buffer that you have: 100 mL = 0,1 liters, and 0,1 liters of 0,1 M TRIS buffer is 0,01 moles. Since the buffer is at its p*K*a, there are equal concentrations of the acid and basic form, so the amount of TRIS is 0,005 moles, and the amount of TRIS-H$^+$ is 0,005 moles.
If you then add 3 mL of 1 M HCl, you will be adding 0,003 moles of H$^+$. This will react as shown:
TRIS + H$^+$ → TRIS-H$^+$ until you run out of something, which will be the H$^+$ since
it is the limiting reagent. The new amounts can be calculated as shown below:
TRIS-H$^+$ = 0,005 moles + 0,003 moles = 0,008 moles
TRIS = 0,005 moles - 0,003 moles = 0,002 moles
Now plug these values into the Henderson-Hasselbalch equation:
pH = 8,3 + log ([TRIS]/[TRIS-H$^+$]) = 8,3 + log(0,002/0,008) pH = 7.70
35. First calculate the moles of buffer that you have: (we are going to do some rounding off) 100 mL = 0,1 liters, and 0,1 liters of 0,1 M TRIS buffer is 0,01 moles. Since the buffer is at pH 7,70, we saw in exercise 25 that the amount of TRIS is 0,002 moles, and the amount of TRIS-H$^+$ is 0,008 moles.
If you then add 3 mL of 1 M HCl, you will be adding 0,003 moles of H$^+$. This will react as shown:
TRIS + H$^+$ → TRIS-H$^+$ until you run out of something, which will be the TRIS
since it is the limiting reagent. All the TRIS is converted to TRIS-H$^+$:
TRIS-H$^+$ = 0,01 moles
TRIS = ~0 moles
We have used up the buffer capacity of the TRIS. We now have 0,003 moles of H$^+$ in
approximately 0,1 liters of solution. This is approximately 0,3M H$^+$.
pH = - log 0,3
pH = 0,52
36. [H+] = [A-] for pure acid, thus Ka =[H+]^2/[HA]
[H+]^2 = Ka x [HA] -2 log [H+] = p*K*a - log [HA]
pH = 1/2 {p*K*a - log [HA]}
37. Use the Henderson-Hasselbalch equation. [Acetate ion]/[acetic acid] = 2,3/1
38. A substance with a p*K*a of 3.9 has a buffer range of 2.9 to 4.9. It will not buffer effectively at pH 7,5.
39. Use the Henderson-Hasselbalch equation. The ratio of A$^-$/HA would be 3981 to 1.
40. In all cases the suitable buffer range covers a pH range of p*K*a +/— 1 pH units.
Lactic acid (p*K*a = 3,86) and its sodium salt, pH 2,86-4,86
Acetic acid (p*K*a = 4.76) and its sodium salt, pH 3.76-5.76
TRIS (see Table 3,4, p*K*a, = 8,3) in its protonated form and its free amine form,
pH 7.3-9.3
HEPES (see Table 3,4, p*K*a, = 7.55) in its zwitterionic and its anionic form, pH 6.55-8.55
41. Several of the buffers would be suitable, namely TES, HEPES, MOPS, and PIPES, however the best buffer would be MOPS as its p*K*a of 7.2 is closest to the desired pH of 7.3.
42. The solution is called 0,0500 molar, even though the concentration of neither the free base nor the conjugate acid is 0,0500 M. Why is 0,0500M the correct concentration to report? Buffer concentrations are typically reported to be the sum of the two ionic forms.)
43. At the equivalence point of the titration a small amount of acetic acid remains because of the equilibrium $CH_3COOH \rightarrow H^+ + CH_3COO^-$. There is a small, but nonzero, amount of acetic acid left.
44. Buffering capacity is based upon the amounts of the acid and base forms present in the buffer solution. A solution with a high buffering capacity can react with large amount of added acid or base without drastic changes in pH. A solution with a low buffering capacity can react with only comparatively small amounts of acid or base before showing changes in pH. The more concentrated the buffer, the higher is its buffering capacity. The first buffer listed here has 10 times less buffering capacity than the second, which in turn has 10 times less buffering capacity than the third. All three buffers have the same pH, since they all have the same relative amounts of the acid and base form.
45. It would be more effective to start with the HEPES base. You want a buffer at a pH above the p*K*a, which means that the base from will predominate when you have finished preparing it. It is easier to convert some of the base form to the acid form than most of the acid form to the base form.
46. In a buffer with the pH above the p*K*a, the base form predominates. This would be useful as a buffer for a reaction that produces H$^+$ because there will be plenty of the base form to react with the hydrogen ion produced.
47. Zwitterions tend not to interfere with biochemical reactions.
48. It is useful to have a buffer that will maintain a stable pH even if assay conditions change. Dilution is one such possible change.
49. It is useful to have a buffer that will maintain a stable pH even if assay conditions change. Temperature variation is one such possible change.
50. The only zwitterion is $^+H_3N{-}CH_2{-}COO^-$.

Chapter 3 Answers

3,1 What Are Amino Acids, and What is Their Three-Dimensional Structure?

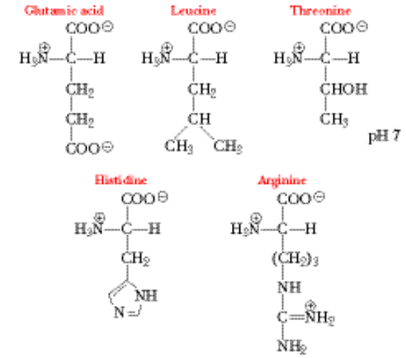
1. D- and L-amino acids have different stereochemistry around the a-carbon. Peptides that contain D-amino acids are found in bacterial cell walls and in some antibiotics.

3,2 What Are the Structures and Properties of the Individual Amino Acids?

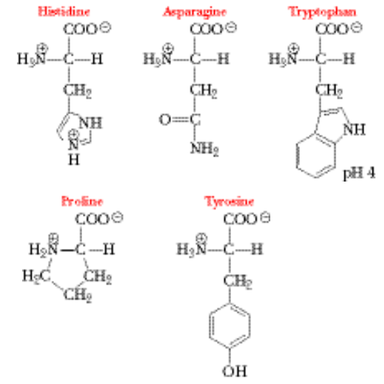
2. Proline is technically not an amino acid. Glycine contains no chiral carbon atoms.
3. An amino acid in which the R group contains the following: a hydroxyl group (serine, threonine, tyrosine) a sulfur atom (cysteine, methionine) a second chiral carbon atom (isoleucine, threonine) an amino group (lysine) an amide group (asparagine, glutamine) an acid group (aspartate, glutamate) an aromatic ring (phenylalanine, tyrosine, tryptophan) a branched side chain (leucine, valine)
4. In the peptide, Val-Met-Ser-Ile-Phe-Arg-Cys-Tyr-Leu, the polar amino acids are Ser, Arg, Cys, and Tyr; the aromatic amino acids are Phe and Tyr; and the sulfur-containing amino acids are Met and Cys.
5. In the peptide, Glu-Thr-Val-Asp-Ile-Ser-Ala, the nonpolar amino acids are Val, Ile, and Ala; the acidic amino acids are Glu and Asp.
6. Amino acids other than the usual 20 are produced by modification of one of the common amino acids. See Figure 3.5 for the structures of some modified amino acids. Hydroxyproline and hydroxylysine are found in collagen; thyroxine is found in thyroglobulin.
7. Tyrosine, tryptophan, and their derivatives.
8. A monoamine oxidase is an enzyme that degrades compounds with an amino group, including neurotransmitters; consequently, they can control a person's mental state.
9. The high concentration of tryptophan in milk protein may mildly elevate the levels of serotonin, which relaxes the brain.
10. The tryptophan in milk might make you sleepy, whereas the tyramine in cheese should pep you up.
11. The amino acids thyroxine and hydroxyproline occur in very few proteins. The genetic code does not include them, so they are produced by modification of tyrosine and proline, respectively.

3,3 Do Amino Acids Have Specific Acid-Base Properties?

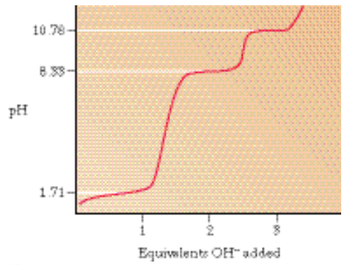
12.



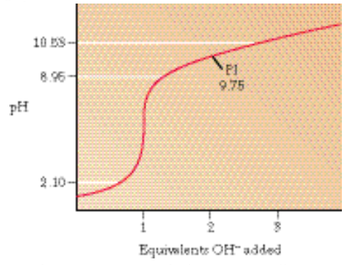
13.



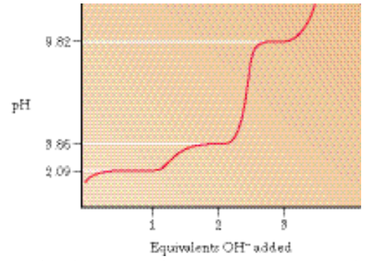
14. Histidine: imidazole is deprotonated, -amino group is predominantly deprotonated, Asparagine: -amino group is deprotonated, Tryptophan: -amino group is predominantly deprotonated, Proline: -amino group is partially deprotonated, Tyrosine: -amino group is predominantly deprotonated, phenolic hydroxyl is approximately a 50 - 50 mixture of protonated and deprotonated forms.
15. Isoelectric point for glutamic acid, 3,25; serine, 5,7; histidine, 7,58; lysine, 9,75; tyrosine, 5,65; arginine, 10,75.
16. Cysteine will have no net charge at pH 5.02 = (1,71 + 8,33)/2 (see titration curve below).



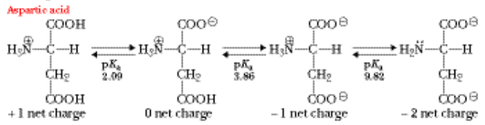
17.

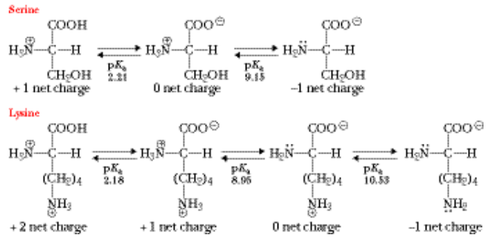


18. The conjugate acid - base pair will act as a buffer in the pH range 1.09 - 3.09.



19. They have a net charge at pH extremes, and the molecules tend to repel each other. When the molecular charge is zero, the amino acids can aggregate more easily.
20. The ionic dissociation reactions of the following amino acids: aspartic acid, valine, histidine, serine, and lysine:





21. The pKa for the ionization of the thiol group of cysteine is 8.33, so this amino acid could serve as a buffer in the -SH and -S- forms over the pH range 7.33-9.33. The - amino groups of asparagine and lysine have pKa values of 8.80 and 8.95, respectively; these are also possible buffers, but they are both near the end of their buffer ranges.
22. At pH 4, the -carboxyl group is deprotonated to a carboxylate, the side chain carboxyl is still more than 50% protonated, and both amino groups are protonated. At pH 7, both the -carboxyl group and the side chain carboxyl group are deprotonated to a carboxylate, and both amino groups are protonated. At pH 10, both the -carboxyl group and the side chain carboxyl group are deprotonated to a carboxylate, one of the amino groups is primarily deprotonated, and the other amino group is a mixture of the protonated and deprotonated forms.
23. The pI refers to the form in which both carboxyl groups are deprotonated and both amino groups protonated at pH 6.96.
23. The pI refers to the form in which both carboxyl groups are deprotonated and both amino groups protonated at pH 6.96.
24. At pH 1 the charged groups are the N-terminal NH_3^+ on valine and the protonated guanidino group on arginine, giving zero net charge. The charged groups at pH 7 are the same as pH 1 with the addition of the carboxylate group on the C-terminal leucine, giving a net charge of -1.
25. Both peptides, Phe-Glu-Ser-Met and Val-Trp-Cys-Leu, have a charge of -1 at pH 1 because of the protonated N-terminal amino group. At pH 7, the peptide on the right has no net charge because of the protonated N-terminal amino group and the ionized Cterminal carboxylate negative charge. The peptide on the left has a net charge of -1 at pH 7 because of the side-chain carboxylate group on the glutamate in addition to the charges on the N-terminal and C-terminal groups.
26. (a) Lysine because of the side chain amino group; (b) Serine because of the lack of a side chain carboxyl; (c) Histidine, because of the presence of a titratable group in the side chain; (d) Aspartate because of the carboxyl group in the side chain.
27. Glycine is frequently used as the basis of a buffer in the acid range near the pK of its carboxyl group. The useful buffer range is from pH 1.3 to 3.3.

3.4 What Is the Peptide Bond?

28. See Figure 3.9.
29. The resonance structures contribute to the planar arrangement by giving the C-N bond partial double bond character.
30. In all cases, the yield is 0.95n. For 10 residues, that means 60% yield; for 50 residues, 8%; and for 100 residues, 0.6%. These are not satisfactory yields. Enzyme specificity gets around the problem.
31. The two peptides differ in amino acid sequence but not in composition.
32. The titration curves of the two peptides will have the same general shape. The pKa values of the -amino and - carboxyl groups will differ. Very careful work will show slight differences in side chain pKa values because of the different distances to the charged groups at the ends of the peptide. Such changes are particularly marked in proteins.
33. Asp-Leu-Phe Leu-Asp-Phe Phe-Asp-Leu Asp-Phe-Leu Leu-Phe-Asp Phe-Leu-Asp
34. DLF LDF FDL DFL LFD FLD
35. You would get 20^{100} ~1.27 × 10^{130} molecules, which is about 10^{84} Earth volumes. The same calculation for a pentapeptide gives more comprehensible results.
36. They are relatively stable because they are zwitterions. They typically have high melting points.
37. With very little doubt, no. Compare predicting the properties of water from those of hydrogen and oxygen, in either atomic or molecular form. If you knew the properties of the protein, you might be able to do the reverse to some extent.
38. These two peptides differ chemically. The open chain has a free C-terminal and Nterminal, but the cyclic peptide has only peptide bonds.
39. Both the C-terminal and the N-terminals of the open-chain peptide can be charged at appropriate pH values, which is not the case with the cyclic peptide. This can provide a basis for separation by electrophoresis.
40. Carbohydrates are not a source of the nitrogen needed for biosynthesis of amino acids.
41. Suggest that your friend show the carboxyl group as a charged carboxylate ($-COO^-$) and the amino group in its charged form ($-NH_3^+$).
42. There are very few side chains that have functional groups to form crosslinks.
43. There will be many more possible conformations because of free rotation around the peptide bond.
44. There would be no possibility of disulfide crosslinks within or between peptide chains, giving more possible conformations. There would not be the possibility of oxidation - reduction reactions involving sulfhydryl and disulfide groups.
45. The big difference would be the loss of stereospecificity in the conformation of any peptide or protein. This would have drastic consequences for the kinds of reactions of the protein.

3.5 Are Small Peptides Physiologically Active?

46. Oxytocin has an isoleucine at position 3 and a leucine at position 8; it stimulates smooth muscle contraction in the uterus during labor and in the mammary glands during lactation. Vasopressin has a phenylalanine at position 3 and an arginine at position 8; it stimulates resorption of water by the kidneys, thus raising blood pressure.
47. The reduced form of glutathione consists of three amino acids with a sulfhydryl group; the oxidized form consists of six amino acids and can be considered the result of linking two molecules of reduced glutathione by a disulfide bridge.
48. Enkephalins are pentapeptides (Y-G-G-F-L, leucine enkephalin, and Y-G-G-F-M, methionine enkephalin), which are naturally occurring analgesics.
49. In most cases, D-amino acids are toxic. They occur in nature in antibiotics and bacterial cell walls.
50. The different stereochemistry of the two peptides leads to different binding with taste receptors and to the sweet taste for one and the bitter taste for the other.

Chapter 4 Answers

4.1 How Does the Structure of Proteins Determine Their Function?

1. (a)(2); (b)(4); (c)(1); (d)(3).
2. When a protein is denatured, the interactions that determine secondary, tertiary, and any quaternary structures are overcome by the presence of the denaturing agent. Only the primary structure remains intact.
3. The random portions of a protein do not contain structural motifs that are repeated within the protein, such as a-helix or b-pleated sheet, but three-dimensional features of these parts of the protein are repeated from one molecule to another. Thus, the term "random" is somewhat of a misnomer.

4.2 What Is the Primary Structure of Proteins?

4. When a protein is covalently modified, its primary structure is changed. The primary structure determines the final three-dimensional structure of the protein. The modification disrupts the folding process.
5. (a) Serine has a small side chain that can fit in any relatively polar environment. (b) Tryptophan has the largest side chain of any of the common amino acids, and it tends to require a nonpolar environment. (c) Lysine and arginine are both basic amino acids; exchanging one for the other would not affect the side chain pKa, in a significant way. Similar reasoning applies to the substitution of a nonpolar isoleucine for a nonpolar leucine.
6. Glycine is frequently a conserved residue because its side chain is so small as to fit in spaces that will not accommodate larger ones.
7. When alanine is replaced by isoleucine, there is not enough room in the native conformation for the larger side chain of the isoleucine. Consequently, there is a great enough change in the conformation of the protein that it loses activity. When glycine is substituted in turn for isoleucine, the presence of the smaller side chain leads to resumption of the active conformation.
8. Meat consists largely of animal proteins and fat. The temperatures

involved in cooking meat are usually more than enough to denature the protein part of the meat.

9. Prion diseases have been linked to the immune system. It is believed that the prion proteins travel in the lymph system bound to lymphocytes and eventually arrive at the nervous tissue where they begin to transform the normal prion protein into the abnormal one.
10. While there are strong genetic predispositions to acquire scrapie, that alone will not cause the disease. The disease must be started by ingesting a prion that already has the altered conformation, PrP^{Sc}.
11. The protein efficiency ratio is an arbitrary measurement of the essential amino acid content of a given type of protein.
12. Eggs have the highest PER.
13. The amino acids that must be consumed in the diet because the body cannot synthesize them in sufficient quantities.
14. Reasons for creating genetically modified foods include increasing their protein content, increasing their shelf life, increasing their resistance to insects or other pests, and decreasing the need for pesticides to grow them.

4.3 What Is the Secondary Structure of Proteins?

15. Shape, solubility, and type of biological function (static, structural versus dynamic, catalytic).
16. The angles of the amide planes as they rotate about the a-carbon. The angles are both defined as zero when the two planes would be overlapping such that the carbonyl group of one contacts the N-H of the other.
17. A β-bulge is a common nonrepetitive irregularity found in antiparallel β-sheets. A misalignment occurs between strands of the β-sheet causing one side to bow outward.
18. A reverse turn is a region of a polypeptide where the direction changes by about 180°. There are two kinds—those that contain proline and those that do not. See Figure 4.6 for examples.
19. The ahelix is not fully extended, and its hydrogen bonds are parallel to the protein fiber. The β-pleated sheet structure is almost fully extended, and its hydrogen bonds are perpendicular to the protein fiber.
20. The aaunit, the bab-unit, the b-meander, the Greek key, the β-barrel.
21. The geometry of the proline residue is such that it does not fit into the a-helix, but it does fit exactly for a reverse turn. See Figure 4.10(c).
22. Glycine is the only residue small enough to fit at crucial points in the collagen triple helix.
23. The principal component of wool is the protein keratin, which is a classic example of α-helical structure. The principal component of silk is the protein fibroin, which is a classic example of β-pleated sheet structure. The statement is somewhat of an oversimplification, but it is fundamentally valid.
24. Wool, which consists largely of the protein keratin, shrinks because of its α-helical conformation. It can stretch and then shrink. Silk consists largely of the protein fibroin, which has the fully extended β-sheet conformation, with far less tendency to stretch or shrink.

4.4 What Can We Say About the Thermodynamics of Protein Folding?

25. (1) Backbone H-bonds, involving the CO and NH groups of the peptide chain; (2) side chain H-bonds, involving any possible hydrogen bond donor or acceptors on the side chains; (3) hydrophobic interactions, involving the nonpolar groups on the protein; (4) electrostatic interactions, involving any charged groups on the protein; (5) metal ligation, involving coordination bonds between side chains and a metal ion.
26. Stabilization of any conformation depends on lowering of energy, regardless of the kind of interaction involved. With small molecule interactions, the main component of energy changes is enthalpic, with relatively small entropy changes. In the case of molecules as large as proteins, the number of possible conformations makes the entropic part of energy changes much more important. This is particularly true in the case of hydrophobic interactions.

4.5 What Is the Tertiary Structure of Proteins?

27. See Figure 4.2 for a hydrogen bond that is part of the α helix (secondary structure). See Figure 4.13 for a hydrogen bond that is part of tertiary structure (side chain hydrogen bonding).
28. See Figure 4.13 for electrostatic interactions, such as might be seen between the side chains of lysine and aspartate.
29. See Figure 4.13 for an example of a disulfide bond.
30. See Figure 4.13 for an example of hydrophobic bonds.
31. A chaperone is a protein that aids another protein in folding correctly and keeps it from associating with other proteins before it has reached its final, mature form.
32. Configuration refers to the position of groups due to covalent bonding. Examples include *cis* and *trans* isomers and optical isomers. Conformation refers to the positioning of groups in space due to rotation around single bonds. An example is the difference between the eclipsed and staggered conformations of ethane.
33. Five possible features limit possible protein configurations and conformations. (1) Although any one of 20 amino acids is possible at each position, only one is used, as dictated by the gene that codes for that protein. (2) Either a D- or an L-amino acid could be used at each position (except for glycine), but only L-amino acids are used. (3) The peptide group is planar so that only *cis* and *trans* arrangements are observed. The *trans* form is more stable and is the one usually found in proteins. (4) The angles ϕ and ψ can theoretically take on any value from 0° to 360°, but some angles are not possible because of steric hindrance; angles that are sterically allowed may not have stabilizing interactions, such as those in the α-helix. (5) The primary structure determines an optimum tertiary structure, according to the "second half of the genetic code."
34. Technically, collagen has quaternary structure because it has multiple polypeptide chains. However, most discussions of quaternary structure involve subunits of globular proteins, not fibrous ones like collagen. Many scientists consider the collagen triple helix to be an example of a secondary structure.

4.6 Can We Predict Folding from Sequence?

35. This level of sequence homology is marginal for use of comparative modeling. It is best to try that method, but then to compare the results with those obtained from the fold recognition approach.
36. See the Protein Data Bank.
37. A prion is a potentially infectious protein found in multiple forms in mammals, often concentrated in nervous tissue. There is a normal form and an abnormal form that tends to form plaques destroying the nervous tissue. They have been found to be transmissible across species.
38. A series of encephalopathies have been found to be caused by the abnormal prion protein. In cows, the disease caused by prions is called bovine spongiform encephalopathy or more commonly mad cow disease. In sheep the disease is called scrapie. In humans it is called Jakob-Creutzfeld disease.
39. The normal form of the prion protein has a higher α-helix content compared to bsheet. The abnormal one has an increased β-sheet content.

4.7 What Is the Quaternary Structure of Proteins?

40. Similarities—both contain heme group; both are oxygen binding; secondary structure is primarily -helix. Differences—hemoglobin is a tetramer, while myoglobin is a monomer; oxygen binding to hemoglobin is cooperative, but noncooperative to myoglobin.
41. The crucial residues are histidines in both proteins.
42. Myoglobin's highest level of organization is tertiary. Hemoglobin's is quaternary.
43. The function of hemoglobin is oxygen transport; its sigmoidal binding curve reflects the fact that it can bind easily to oxygen at comparatively high pressures and release oxygen at lower pressures. The function of myoglobin is oxygen storage; as a result, it is easily saturated with oxygen at low pressures, as shown by its hyperbolic binding curve.
44. In the presence of H^+ and CO_2, both of which bind to hemoglobin, the oxygenbinding capacity of hemoglobin decreases.
45. In the absence of 2,3-*bis*phosphoglycerate, the binding of oxygen by hemoglobin resembles that of myoglobin, characterized by lack of cooperativity. 2,3- *Bis*phosphoglycerate binds at the center of the hemoglobin molecule, increases cooperativity, stabilizes the deoxy conformation of hemoglobin, and modulates the binding of oxygen so that it can easily be released in the capillaries.
46. Fetal hemoglobin binds oxygen more strongly than adult hemoglobin. See Figure 4.25.
47. Histidine 143 in a β-chain is replaced by a serine in a γ-chain.
48. Deoxygenated hemoglobin is a weaker acid (has a higher p*K*a) than oxygenated hemoglobin. In other words, deoxygenated hemoglobin binds more strongly to H^+ than does oxygenated hemoglobin. The binding of H^+ (and of CO_2) to hemoglobin favors the change in quaternary structure to the deoxygenated form of hemoglobin.
49. The primary flaw in your friend's reasoning is a reversal of the definition of pH, which is pH = -log [H^+]. If the release or binding of hydrogen ion by hemoglobin were the primary factor in the Bohr effect, the pH changes would be the opposite of those actually observed. The response of hemoglobin to changes in pH is the central point. When the pH increases, the hydrogen ion concentration decreases, and vice versa.
50. The change of a histidine to a serine in the γ-chain removes a positively charged amino acid that could have interacted with BPG. Thus there are fewer salt bridges to break, so binding is easier than it is in a β-chain.
51. Persons with sickle-cell trait have some abnormal hemoglobin. At high altitudes, there is less oxygen, and the concentration of the deoxy form of the abnormal hemoglobin increases. Less oxygen can be bound, causing the observed breathing difficulties.
52. In fetal hemoglobin, the subunit composition is $\alpha_2\gamma_2$ with replacement of the β-chains by the γ-chains. The sickle-cell mutation affects the β-chain, so the fetus homozygous for HbS has normal fetal hemoglobin.
53. The relative oxygen affinities allow oxygen to be taken by the fetal cells from the maternal Hb.
54. Because people with sickle-cell disease are chronically anemic, some cells with fetal Hb are produced to help overcome the impaired oxygen delivery system.
55. The crystalline form changed because oxygen entered under the cover slip, transforming deoxyhemoglobin to oxyhemoglobin.

Chapter 5 Answers

5.1 How Do We Extract Pure Proteins from Cells?

1. Blender, Potter-Elvejhem, sonicator
2. If you needed to maintain the structural integrity of the subcellular organelles, a Potter- Elvejhem would be better because it is more gentle. The tissue, such as liver, must be soft enough to use with this device.
3. Salting out is a process whereby a highly ionic salt is used to reduce the solubility of a protein until it comes out of solution and can be centrifuged. The salt forms ion · dipole bonds with the water in the solution, which leaves less water available to hydrate the protein. Nonpolar side chains begin to interact between protein molecules, and they become insoluble.
4. Their amino acid content and arrangements make some proteins more soluble than others. A protein with more highly polar amino acids on the surface will be more soluble than one with more hydrophobic ones on the surface.
5. Homogenize the liver cells using a Potter-Elvejhem homogenizer. Spin the homogenate at 500 X g to sediment the unbroken cells and nuclei. Centrifuge the supernatant at 15,000 X g, and collect the pellet, which contains the mitochondria.
6. No, peroxisomes and mitochondria have overlapping sedimentation characteristics. Other techniques, such as sucrose gradient centrifugation, would have to be used to separate the two organelles.

7. If the protein were cytosolic, once the cells were broken open, you could centrifuge at 100,000 X g, and all the organelles would be in the pellet. Your enzyme would be in the supernatant, along with all the other cytosolic ones.
8. Isolate the mitochondria via differential or sucrose gradient centrifugation. Use another homogenization technique combined with a strong detergent to release the enzyme from the membrane.
9. Tables exist to tell you how many grams of ammonium sulfate (AS) to add to get a certain percent saturation. A good plan would be to take the homogenate and add enough AS to yield a 20% saturated solution. Let the sample sit for 15 minutes on ice and then centrifuge. Separate the supernatant from the precipitate. Assay both for the protein you are working with. Add more AS to the supernatant to arrive at a 40% saturated solution, and repeat the process. In this way, you will find out what the percent saturation in AS needs to be to precipitate it.
10. Reasonably harsh homogenization would be able to liberate the soluble protein X from the peroxisomes, which are fragile. Centrifugation at 15,000 X g would sediment the mitochondria (broken or intact). The supernatant would then have protein X but no protein Y. Freeze thaw techniques and sonication would accomplish the same thing, or the mitochondria and the peroxisomes could be separated initially by sucrose gradient centrifugation.

5.2 What Is Column Chromatography?

11. (a) Size, (b) specific ligand binding ability, (c) net charge
12. The largest proteins elute first; the smallest elute last. Larger proteins are excluded from the interior of the gel bead so they have less available column space to travel. Essentially, they travel a shorter distance and elute first.
13. A compound can be eluted by raising the salt concentration or by adding a mobile ligand that has a higher affinity for the bound protein than the stationary resin ligand does. Salt is cheaper but less specific. A specific ligand may be more specific but is often expensive.
14. A compound can be eluted by raising the salt concentration or by changing the pH. Salt is cheap, but it might not be as specific for a particular protein. Changing the pH may be more specific for a tight pI range, but extremes of pH may also denature the protein.
15. Raising the salt concentration is relatively safe. Most proteins will elute this way and still be active in the case of an enzyme. The salt can be removed later via dialysis if necessary. Changing the pH enough to remove the charge can cause the proteins to become denatured. Many proteins are not soluble at the isoelectric points.
16. The basis of most resins is agarose, cellulose, dextran, or polyacrylamide.
17. See Figure 5.7.
18. Within the fractionation range of a gel filtration column, molecules will elute with a linear relationship of log MW versus their elution volumes. A series of standards can be run to standardize the column, and then an unknown can be determined by measuring its elution volume and comparing to a standard curve.
19. Both proteins would elute in the void volume together and would not be separated.
20. Yes, the β-amylase would come out in the void volume, but the BSA would be included in the column bead and would elute more slowly.
21. Set up an anion exchange column, such as Q-Sepharose (quaternary amine). Run the column at pH 8.5, a pH at which the protein X has a net negative charge. Put a homogenate containing protein X on the column and wash with the starting buffer. Protein X will bind to the column. Then elute by running a salt gradient.
22. Use a cation exchange column, like CM-Sepharose, and run it at pH 6. Protein X will have a positive charge and will stick to the column.
23. With a quaternary amine, the column resin always has a net positive charge, and you don't have to worry about the pH of your buffer altering the form of the column. With a tertiary amine, there is a dissociable hydrogen, and the resin may be positive or neutrally charged depending on the buffer pH.
24. The easiest way would be to use a sucrose gradient to separate the mitochondria from the peroxisomes first. Then break open the mitochondria via harsh homogenization or sonication. Centrifuge the mitochondria. The pellet would contain protein B, while the supernatant contained protein A. Contaminants could still exist, but they can be cleaned away by running gel filtration, on Sephadex G-75, which would separate C from A and B, and then ion exchange chromatography on Q-Sepharose at pH 7.5. Protein B would be neutral and would elute, while protein A would stick to the column.
25. Glutamic acid will be eluted first because the column pH is close to its pI. Leucine and lysine will be positively charged and will stick to the column. To elute leucine, raise the pH to around 6. To elute lysine, raise the pH to around 11.
26. A nonpolar mobile solvent will move the nonpolar amino acids fastest, so phenylalanine will be the first to elute, followed by glycine and then glutamic acid.
27. The nonpolar amino acids will stick the most to the stationary phase, so glutamic acid will move the fastest, followed by glycine and then phenylalanine.
28. A protein solution from an ammonium sulfate preparation is passed over a gel filtration column where the proteins of interest will elute in the void volume. The salt, being very small, will move through the column slowly. In this way, the proteins will leave the salt behind and exit the column without it.

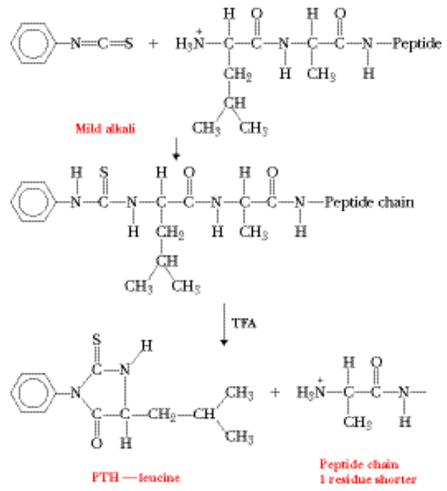
5.3 What Is Electrophoresis?

29. Size, shape, and charge.
30. Agarose and polyacrylamide.
31. Polyacrylamide.
32. DNA is the molecule most often separated on agarose electrophoresis, although proteins can also be separated.
33. Those with the highest charge/mass ratio would move the fastest. There are three variables to consider, and most electrophoreses are done in a way to eliminate two of the variables so that the separation is by size or by charge, but not by both.

34. Sodium dodecyl sulfate polyacrylamide gel electrophoresis. With SDS-PAGE, the charge and shape differences of proteins are eliminated so that the only parameter determining the migration is the size of the protein.
35. SDS binds to the protein in a constant ratio of 1.4 g SDS per gram of protein. It coats the protein with negative charges and puts it into a random coil shape. Thus, charge and shape are eliminated.
36. In a polyacrylamide gel used for gel filtration chromatography, the larger proteins can travel around the beads, thereby having a shorter path to travel and eluting faster. With electrophoresis, the proteins are forced to go through the matrix, so they travel more slowly if they are bigger because there is more friction.
37. The MW is 37,000 daltons.

5.4 How Do We Determine the Primary Structure of a Protein?

38. The Edman degradation will give the identity of the N-terminus in its first cycle, so doing a separate experiment is not necessary.
39. It might tell you if the protein was pure or if there were subunits.
40.



41. The amount of Edman reagent must exactly match the amount of N-termini in the first reaction. If there is too little Edman reagent, some of the N-termini will not react. If there is too much, some of the second amino acid will react. In either case, there will be a small amount of contaminating PTH derivatives. This error grows with the number of cycles run until the point that two amino acids are released in equal amounts, and you could not tell which one was supposed to be the correct one.
42. In the first cycle, the first and second amino acids from the N-terminal end would be reacted and released as PTH derivatives. You would get a double signal and not know which one was the true N-terminus.
43. Val-Leu-Gly-Met-Ser-Arg-Asn-Thr-Trp-Met-Ile-Lys-Gly-Tyr-Met-Gln-Phe
44. Met-Val-Ser-Thr-Lys-Leu-Phe-Asn-Glu-Ser-Arg-Val-Ile-Trp-Thr-Leu-Met-Ile
45. It is possible that your protein is not pure and needs additional purification steps to arrive at a single polypeptide. It is also possible that the protein has subunits, so multiple polypeptide chains could be yielding the contradictory results.
46. There are two fragments that have C-termini that are not lysine or arginine, which is what trypsin is specific for. Normally there would be only one fragment ending with an amino acid that was not Arg or Lys, and we would immediately know that it was the Cterminus. Histidine is a basic protein, although it is usually neutral and therefore does not react with trypsin. It is possible that in the pH environment of the reaction, the histidine was positively charged and was recognized by trypsin.
47. It would tell you a relative concentration of the various amino acids. This is important because it would help you plan your sequencing experiment better. For example, if you had a protein whose composition showed no aromatic amino acids, it would be a waste of time to use a chymotrypsin digestion.
48. Cyanogen bromide would be useless as there is no methionine. Trypsin would be little better as the protein is 35% basic residues. Trypsin would shred the protein into over 30 pieces, which would be very hard to analyze.
49. Chymotrypsin would be a good choice. There are more than four residues of aromatic amino acids. The 100 amino acid protein would be cut four times, possibly yielding nice fragments in the 20 - 30 amino acid length, which can be sequenced effectively by the Edman degradation.
50. It would work best if the basic residues were spread out in the protein. In that way, fragments in the proper size range would be generated. If all four of the basic residues were in the first ten amino acids, there would be one long fragment that could not be sequenced.

Chapter 6 Answers

6.1 What Makes Enzymes Such Effective Biological Catalysts?

1. Enzymes are many orders of magnitude more effective as catalysts than are nonenzymatic catalysts.
2. The majority of enzymes are proteins, but some catalytic RNAs (ribozymes) are known.
3. About 3 seconds (1 year X 1 event/10^7 events X 365 days/year X 24 hours/day X 3600 seconds/hour = 3.15 seconds).
4. Enzymes hold the substrates in favorable spatial positions, and they bind effectively to the transition state to stabilize it. Note that all catalysts lower the activation energy, so this is not a particular enzyme function.

6.2 What Is the Difference between the Kinetic and Thermodynamic Aspects of Reactions?

5. The reaction of glucose with oxygen is thermodynamically favored,

as shown by the negative free energy change. The fact that glucose can be maintained in an oxygen atmosphere is a reflection of the kinetic aspects of the reaction, requiring overcoming an activation-energy barrier.

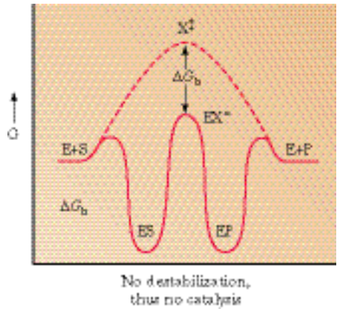
6. First question, most probably; local concentrations (mass action concepts) could easily dictate the direction. Second question, probably not; local concentrations would seldom be sufficient to overcome a relatively large ΔG° of -5.3 kcal in the reverse reaction. (See, however, the aldolase reaction in glycolysis.)
7. Heating a protein denatures it. Enzymatic activity depends on the correct threedimensional structure of the protein. The presence of bound substrate can make the protein harder to denature.
8. The results do not prove that the mechanism is correct because results from different experiments could contradict the proposed mechanism. In that case, the mechanism would have to be modified to accommodate the new experimental results.
9. The presence of a catalyst affects the rate of a reaction. The standard free energy change is a thermodynamic property that does not depend on the reaction rate. Consequently, the presence of the catalyst has no effect.
10. The presence of a catalyst lowers the activation energy of a reaction.
11. Enzymes, like all catalysts, increase the rate of the forward and reverse reaction to the same extent.
12. The amount of product obtained in a reaction depends on the equilibrium constant. A catalyst does not affect that.

6.3 How Can We Describe Enzyme Kinetics in Mathematical Terms?

13. The reaction is first order with respect to A, first order with respect to B, and second order overall. The detailed mechanism of the reaction is likely to involve one molecule each of A and B.
14. The easiest way to follow the rate of this reaction is to monitor the decrease in absorbance at 340 nm, reflecting the disappearance of NADH.
15. The use of a pH meter would not be a good way to monitor the rate of the reaction. You are probably running this reaction in a buffer solution to keep the pH relatively constant. If you are not running the reaction in a buffer solution, you run the risk of acid denaturation of the enzyme.
16. Enzymes tend to have fairly sharp pH optimum values. It is necessary to ensure that the pH of the reaction mixture stays at the optimum value. This is especially true for reactions that require or produce hydrogen ions.

6.4 How Do Substrates Bind to Enzymes?

17. In the lock-and-key model, the substrate fits into a comparatively rigid protein that has an active site with a well-defined shape. In the induced-fit model, the enzyme undergoes a conformational change on binding to substrate. The active site takes shape around the substrate.
18.



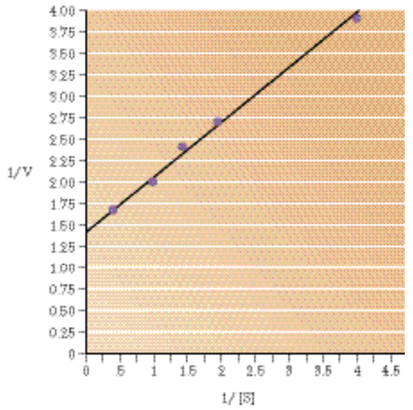
19. The ES complex would be in an "energy trough," with a consequentially large activation energy to the transition state.
20. Amino acids that are far apart in the amino acid sequence can be close to each other in three dimensions because of protein folding. The critical amino acids are in the active site.
21. The overall protein structure is needed to ensure the correct arrangement of amino acids in the active site.
22. The strong inhibition indicates tight binding to the active site. Thus, the compound is very likely to be a transition state analogue.

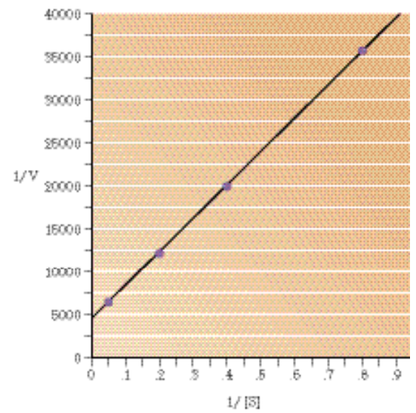
6.5 What Are Some Examples of Enzyme-Catalyzed Reactions?

23. See Figures 6.6 and 6.7.
24. Not all enzymes follow Michaelis - Menten kinetics. The kinetic behavior of allosteric enzymes does not obey the Michaelis - Menten equation.
25. The graph of rate against substrate concentration is sigmoidal for an allosteric enzyme but hyperbolic for an enzyme that obeys the Michaelis - Menten equation.

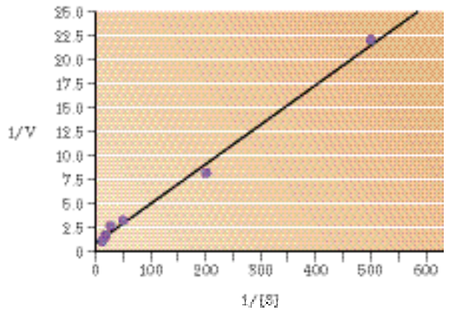
6.6 What Is the Michaelis - Menten Approach to Enzyme Kinetics?

26. The reaction velocity remains the same with increasing enzyme concentration. It is theoretically possible, but highly unlikely, for a reaction to be saturated with enzyme.
27. The steady-state assumption is that the concentration of the enzyme - substrate complex does not change appreciably over the time in which the experiment takes place. The rate of appearance of the complex is set equal to its rate of disappearance, simplifying the equations for enzyme kinetics.
28. Turnover number = $V\text{max}/[E]_t$.
29. Use Equation 6.16. (a) $V = 0.5\ V\text{max}$; (b) $V = 0.33\ V\text{max}$; (c) $V = 0.09\ V\text{max}$; (d) $V = 0.67\ V\text{max}$; (e) $V = 0.91\ V\text{max}$.
30. See graph: $V\text{max} = 0.681\ \text{m}M\,\text{min}^{-1}$, $K_M = 0.421\ M$.

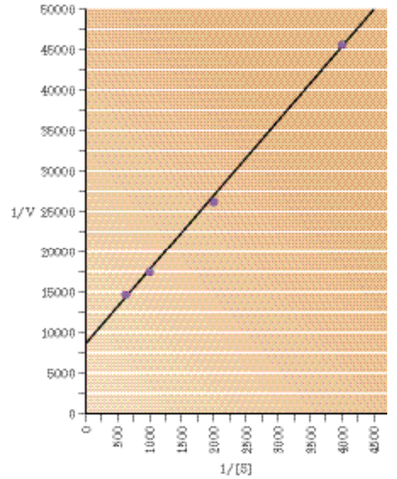




31. See graph: $V\text{max} = 2.5 \times 10^{-4}\ M\,\text{sec}^{-1}$, $K_M = 1.6 \times 10^{8}\ M$.
32. See graph: $K_M = 2.86 \times 10^{-2}\ M$. Concentrations were not determined directly. Absorbance values were used instead as a matter of convenience.



33. See graph: $V\text{max} = 1.32 \times 10^{-3}\ M\,\text{min}^{-1}$, $K_M = 1.23 \times 10\text{-}3\ M$.



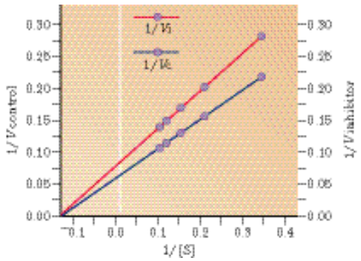
34. The turnover number is 20,43 per minute.
35. The number of moles of enzyme is 1.56×10^{-10}. Turnover number = 10,700 sec^{-1}.
36. The low K_M for the aromatic amino acids indicates that they will be oxidized preferentially.
37. It is easier to detect deviations of individual points from a straight line than from a curve.
38. The assumption that the K_M is an indication of the binding affinity between the substrate and the enzyme is valid when the rate of dissociation of the enzyme - substrate complex to product and enzyme is much smaller than the rate of dissociation of the complex to enzyme and substrate.

6.7 How Do Enzymatic Reactions Respond to Inhibitors?

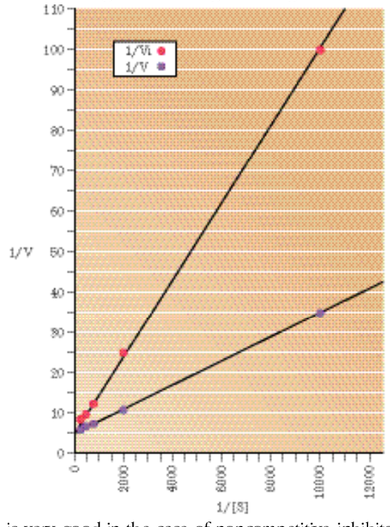
39. In the case of competitive inhibition, the value of K_M increases, while the value of K_M remains unchanged in noncompetitive inhibition.
40. A competitive inhibitor blocks binding, not catalysis.
41. A noncompetitive inhibitor does not change the affinity of the enzyme for its substrate.
42. A competitive inhibitor binds to the active site of an enzyme, preventing binding of the substrate. A noncompetitive inhibitor binds at a site different from the active site, causing a conformational change, which renders the active site less able to bind substrate and convert it to product.
43. Competitive inhibition can be overcome by adding enough substrate, but this is not true for all forms of enzyme inhibition.
44. A Lineweaver - Burk plot is useful because it gives a straight line. It is easier to determine how well points fit to a straight line than to a curve.
45. In a Lineweaver - Burk plot for competitive inhibition, the lines intersect at the y-axis intercept, which is equal to $1/V\text{max}$. In a Lineweaver - Burk plot for noncompetitive inhibition, the lines

intersect at the x-axis intercept, which is equal to - 1/KM.

46. K_M = 7.42 m*M*; *V*max = 15.9 mmol min^{-1}; noncompetitive inhibition.



47. Competitive inhibition, $K_M = 6,5 \times 10^{-4}$. The key point here is that the Vmax is the same within the limits of error. Some of the concentrations are given to one significant figure.



48. It is very good in the case of noncompetitive inhibitors; much of metabolic control depends on feedback inhibition by downstream noncompetitive inhibitors. The question is perhaps moot in the case of competitive inhibitors, which are much less commonly encountered *in vivo*. Some antibiotics, however, are competitive inhibitors (good for the sick person, bad for the bacteria).
49. Both the slope and the intercepts will change. The lines will intersect above the x-axis at negative values of 1/[S].
50. Not all AIDS drugs are enzyme inhibitors, but an important class of such drugs inhibits the HIV protease. You would need to understand the concepts of substrate binding, inhibition, and inhibitor binding.
51. An irreversible inhibitor is bound by covalent bonds. Noncovalent interactions are relatively weak and easily broken.
52. A noncompetitive inhibitor does not bind to the active site of an enzyme. Its structure need bear no relation to that of the substrate

Chapter 7 Answers

7.1 Does the Michaelis - Menten Model Describe the Behavior of Allosteric Enzymes?

1. Allosteric enzymes display sigmoidal kinetics when rates are plotted versus substrate concentration. Michaelis-Menten enzymes exhibit hyperbolic kinetics. Allosteric enzymes usually have multiple subunits, and the binding of substrates or effector molecules to one subunit changes the binding behavior of the other subunits.
2. It is an enzyme used in the early stages of cytidine nucleotide synthesis.
3. ATP acts as a positive effector of ATCase and CTP acts as an inhibitor.
4. The term K_M should be used for enzymes that display Michaelis - Menten kinetics. Thus, it is not used with allosteric enzymes. Technically, competitive and noncompetitive inhibition are also terms that are restricted to Michaelis - Menten enzymes, although the concepts are applicable to any enzyme. An inhibitor that binds to an allosteric enzyme at the same site as the substrate is similar to a classical competitive inhibitor. One that binds at a different site is similar to a noncompetitive inhibitor, but the equations and the graphs characteristic of competitive and noncompetitive inhibition don't work the same way with an allosteric enzyme.
5. A K system is an allosteric enzyme in which the binding of inhibitor alters the apparent substrate concentration needed to reach one-half V_{max}, $S_{0.5}$.
6. A V system is an allosteric enzyme in which the binding of inhibitor changes the V_{max} of the enzyme but not the $S_{0.5}$.
7. Homotropic effects are allosteric interactions that occur when several identical molecules are bound to a protein. The binding of substrate molecules to different sites on an enzyme, such as the binding of aspartate to ATCase, is an example of a homotropic effect. Heterotropic effects are allosteric interactions that occur when different substances (such as inhibitor and substrate) are bound to the protein. In the ATCase reaction, inhibition by CTP and activation by ATP are both heterotropic effects.
8. ATCase is made up of two different types of subunits. One of them is the catalytic subunit, and there are six of them organized into two trimers. The other is the regulatory subunit, which also consists of six protein subunits organized into three dimers.
9. Enzymes that exhibit cooperativity do not show hyperbolic curves of rate versus substrate concentration. Their curves are sigmoidal. The level of cooperativity can be seen by the shape of the sigmoidal curve.
10. Inhibitors make the shape of the curve more sigmoidal.
11. Activators make the shape of the curve less sigmoidal.
12. $K_{0.5}$ is the substrate concentration that leads to half of the maximal velocity. This term is used with allosteric enzymes where the term K_M is not appropriate.
13. A mercury compound was used to separate the subunits of ATCase. When the subunits were separated, one type of subunit retained catalytic activity but was no longer allosteric and was not inhibited by CTP. The other subunit type had no ATCase activity, but it did bind to CTP and ATP.

7.2 What Are the Models for the Behavior of Allosteric Enzymes?

14. In the concerted model, all the subunits in an allosteric enzyme are found in the same form, either the T or the R. They are in equilibrium with each enzyme having a characteristic ratio of the T/R. In the sequential model, the subunits change individually from T to R.
15. The sequential model can explain negative cooperativity because a substrate binding to the T form could induce other subunits to switch to the T form, thereby reducing binding affinity.
16. Greater cooperativity is favored by having a higher ratio of the T/R form. It is also favored by having a higher dissociation constant for the substrate binding to the T form.
17. The L-value is the equilibrium ratio of the T/R form. The c value is the ratio of the dissociation constants for substrate and the two forms of enzyme, such that
c = *K*R/*K*T.
18. Many models are possible. We never really know for sure how the enzyme works, rather we create a model that explains the observed behavior. It is very possible that another model would do so as well.

7.3 How Does Phosphorylation of Specific Residues Regulate Enzyme Activity?

19. A kinase is an enzyme that phosphorylates a protein using a high-energy phosphate, such as ATP, as the phosphate donor.
20. Serine, threonine, and tyrosine are the three most often phosphorylated amino acids in proteins that are acted upon by kinases. Aspartate is another one often phosphorylated.
21. The allosteric effect can be faster because it is based on simple binding equilibrium. For example, if AMP is an allosteric activator of glycogen phosphorylase, the immediate increase in AMP when muscles contract can cause muscle phosphorylase to become more active and provide energy for the contracting muscles. The phosphorylation effect requires the hormone cascade beginning with glucagon or epinephrine. There are many steps before the glycogen phosphorylase is phosphorylated, so the response time is slower. However, the cascade effect produces many more activated phosphorylase molecules, so the effects are longer and stronger.
22. As part of the mechanism, the sodium-potassium ATPase has an aspartate residue that becomes phosphorylated. This phosphorylation alters the conformation of the enzyme and causes it to close on one side of the membrane and open on the other, moving ions in the process.
23. Glycogen phosphorylase is controlled allosterically by several molecules. In the muscle, AMP is an allosteric activator. In the liver, glucose is an allosteric inhibitor. Glycogen phosphorylase also exists in a phosphorylated form and an unphosphorylated form, with the phosphorylated form being more active.

7.4 What Are Zymogens, and How Do They Control Enzyme Activity?

24. The digestive enzymes trypsin and chymotrypsin are classic examples of regulation by zymogens. The blood clotting protein thrombin is another.
25. Trypsin, chymotrypsin, and thrombin are all proteases. Trypsin cleaves peptide bonds where there are amino acids with positively charged side chains (Lys and Arg). Chymotrypsin cleaves peptides at amino acids with aromatic side chains. Thrombin cleaves the protein fibrinogen into fibrin.
26. The zymogen prothrombin is cleaved to give the active enzyme thrombin. The thrombin then cleaves a soluble molecule, fibrinogen, into an insoluble molecule, fibrin. Fibrin is a protein that forms part of the blood clot.
27. Chymotrypsinogen is an inactive zymogen. It is acted upon by trypsin, which cleaves peptides at basic residues, like arginine. When trypsin cleaves between the arginine and the isoleucine, chymotrypsinogen becomes semiactive, forming π-chymotrypsin. This molecule digests itself further, forming the active α-chymotrypsin. As it turns out, the α-amino group of the isoleucine produced by the first cleavage is near the active site of α-chymotrypsin and necessary for its activity.
28. Zymogens are often seen with digestive enzymes that are produced in one tissue and used in another. If the enzyme were active immediately upon production, it would digest other cell proteins where it would cause great damage. By having it produced as a zymogen, it can be safely made and then transported to the digestive tissue, such as the stomach or small intestine, where it can then be activated.
29. This allows for a more rapid response when the hormone is needed. The hormone is already synthesized and usually just requires breaking one or two bonds to make it active. The hormone can be poised and ready to go on demand.

7.5 How Do Active Site Events of an Enzyme Affect the Reaction Mechanism?

30. Serine and histidine are the two most critical amino acids in the active site of chymotrypsin.
31. The initial phase releases the first product and involves an acyl-enzyme intermediate. This step is faster than the second part where water comes into the active site and breaks the enzyme-acyl bond.
32. In the first step of the reaction the serine hydroxyl is the nucleophile that attacks the substrate peptide bond. In the second step, water is the nucleophile that attacks the acylenzyme intermediate.
33. His57 performs a series of steps involving general base catalysis followed by general acid catalysis. In the first phase, it takes a hydrogen from Ser195, acting as a general base. This is followed immediately by an acid catalysis step where it gives the hydrogen to the amide group of the peptide bond that is breaking. A similar scheme takes place in the second phase of the reaction.
34. The first phase is faster for several reasons. The serine at position

195 is a strong nucleophile for the initial nucleophilic attack. It then forms an acyl enzyme intermediate. In the second phase, water is the nucleophile, and it takes time for water to diffuse to the right spot to perform its nucleophilic attack. It is also not as strong a nucleophile as the serine. Therefore, it takes longer for water to perform its nucleophilic attack and break the acyl-enzyme intermediate than it takes for serine to create it.

35. His57 exists in both the protonated and unprotonated form during the chymotrypsin reaction. Its pKa of 6.0 makes this possible in the physiological pH range.
36. Instead of a phenylalanine moiety (similar to the usual substrates of chymotrypsin), use a nitrogen-containing basic group similar to the usual substrates of trypsin.

7.6 What Types of Chemical Reactions are Involved in Enzyme Mechanisms?

37. They act as Lewis acids (electron pair acceptors) and can take part in enzyme catalysis mechanisms of enzymes.
38. False. The mechanisms of enzymatic catalysis are the same as those encountered in organic chemistry, operating in a complex environment.
39. General acid catalysis is the part of an enzyme mechanism where an amino acid or other molecule donates a hydrogen ion to another molecule.
40. S_N1 stands for unimolecular nucleophilic substitution. The unimolecular part means that it obeys first-order kinetics. If the reaction is R:X + Z: → R:Z + X:, with an S_N1 reaction, the rate is dependent on the speed with which the X breaks away from the R. The Z group comes in later and quickly, compared to the breakdown of R:X. S_N2 stands for bimolecular nucleophilic substitution. This happens with the same reaction scheme if the Z attacks the R:X molecule before it breaks down. Thus the concentration of both R:X and Z: are important, and the rate displays second-order kinetics.
41. The S_N1 reaction leads to loss of stereospecificity as the X group leaves before the entering nucleophile. This means that the nucleophile can enter from different angles leading to different isomers.
42. The results do not prove that the mechanism is correct because results from different experiments could contradict the proposed mechanism. In that case, the mechanism would have to be modified to accommodate the new experimental results.

7.7 What Is the Connection between the Active Site and Transition States?

43. A good transition state analogue would have to have a tetrahedral carbon atom where the amide carbonyl group was originally found, since the transition state involves a momentary tetrahedral form. It would also have to have oxygens on the same carbon so that there would be sufficient specificity for the active site.
44. The induced fit model assumes that the enzyme and substrate must both move and change to conform to each other perfectly. Thus, the true fit is not between the enzyme and substrate but between the enzyme and the transition state of the substrate on its way to product. A transition state analogue will fit the enzyme nicely in this model.

Sw.Rs tOPROB Le mSCHAPter 1A|SWeRSL o 1 wha tafe theRs i Chemsfor thisT exT'll AP oly merisive rY larGemOL.CUi e for medoyl i fli ngSm auTeTuflits (moflo fllers)to gether1A p rotTinis aroLYmETfro rTed byti nkingflli flaCiaS toBeth ei A nucle ic acidi s apoLymeTfor meabY-ilking nucle.. tid estoSet le n catal y si sisthePr oES sthatjncr e

asSsther dted fchemicdur eaction SComPa red toth eunc at aly zed-eaction fi0i0 gical cat al ySts arep. Otei n Sinaulos tuic ds tS; theonu Yexcep, tionsfle afe typ eSo rRNA>w hiC hcaNat a l YZe someof theredc le ion Sorth ei r0 Inme ta nOlisthm. Gefet i Coq ei Sthell e ansbyThi chtifC in f0 rmati on f0 rtheStr uCtiraNd ruft ctiOn 0ra lulliy jingt hifig sis pasS ed fr0mOfeg ener a tio n

t0. henetu the Se qUenCeof Pu r ines fldp yrimi d ines inDiACarri es tuse fleti tCo d c RNi st ecodi n fiaterialin. 0meVi ruses>u l2Wha tiSthe Cofm, cal natufeo erim orta ntbiom0 l ecule2) T ecor rect nltchor fln ction. Igrolpsin dc0Poulus co nta inin shdtur uicti ol dlg r0Ubi Sgi e ri nuneforl 0Winsyl is u amino gr0UP CHBCH2NH 2cgr0n-igro Up<ketin—) CHBCOC HBHY dr0 xyl gr0up CHBOH, arhox. Igro UpCHBCOO, car bonylgr ourc al dehyd e>CHBOH2 CHOTHi0l. ro UpC-3SHEstri ifldg eCHBCOOC HCHBBlou bieron, CH3 CHc HCHBAlide iinkage cH3Co NcH3>2ith erCH3OH2..

C,2CHBBl Th e function al

Ig. Olusiftuhe COm pOundsf Olow4 ltfo, er 82s the-onCeptofvi tluism neidtha tUrsali coMPo undSc0uld-niYbeMdeb yliviNgss Ys tenS anawefere yondthe al mpflato, atorynvestisat iOnslw0 hiel' ssln-hesi sSh0wed thUt0 regfli comPou..

dS. Iiie inorganic one S,diamtreq ui reavitali st icCl phundti.. n>b utrathe, tleVoleyedth e lawSof ClemiSturaNd PhysiCsand thus we re sbieCt tOrato. ator'i nVestig ati0n dsubSeq uentl Y;th e

C0..C0twds eX tende dtothe mlCh0lecol plexybuts, il lte Stabie ld isci pli feofbio chemiStry.

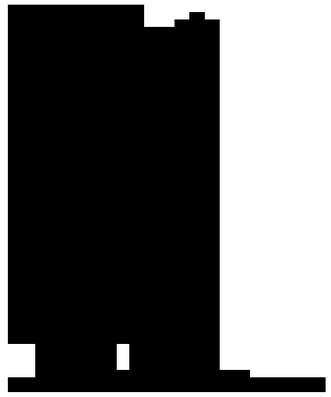
5l. Reay likea no rgani COMP0Und s? ha sthesame m0lecula rstru cUlfe Wh0t heritiSer.. duc eabyali v i ngOrganisMO rmU 6. 7lF-ve:7 ifth et Wocy. Iopro rai n ederivat i vesd re all owed ? T hir-Eendif re rEntialco hOts,l i aldehyd es/ ket on e s> alkl0eZth eP oxidSan dEthers l0 3.

hd. Can i0che mistr Ys a yAb0utpOss. bleorig ins orLife? 9 u t iSg effera uYtetiev edth at carbonl isth eli ke lyba s isf Orallli fef orms? eart hso unato fextrater f estri al l 0l E i Sitee pres dles. 0Uldsive2, kOr2l 6\23>ossibi li ti es! Thu sl9r..

Sidyes W 0 Uldbene

C.SSary to hav eatlelstAvoga dfO Sfl

U.. hr Cl02 X 1023 orPoss iti li ti Es h l 0 the num be ri Sam orb 2×10, 4> Th-Chi St wiceAVog adfo stUll. en 1 2lRAi sca pab le ofb otcoa nsflld



12. The waxy surface coating is a barrier that prevents loss of water.
13. The surface wax keeps produce fresh by preventing loss of water.
14. Cholesterol is not very water soluble, but lecithin is a good natural detergent, which is actually part of lipoproteins that transport the less soluble fats through the blood.
15. The lecithin in the egg yolks serves as an emulsifying agent by forming closed vesicles. The lipids in the butter (frequently triacylglycerols) are retained in the vesicles and do not form a separate phase.
16. The removal of the oil also removes the natural oils and waxes on the feathers. They must regenerate before the birds can be released.

8.2 What Are the Chemical Natures of the Lipid Types?

2. In both types of lipids, glycerol is esterified to carboxylic acids, with three such ester linkages formed in triacylglycerols and two in phosphatidyl ethanolamines. The structural difference comes in the nature of the third ester linkage to glycerol. In phosphatidyl ethanolamines, the third hydroxyl group of glycerol is esterified not to a carboxylic acid, but to phosphoric acid. The phosphoric acid moiety is esterified in turn to ethanolamine. (See Figures 8.2 and 8.5.)
3.

4. Both sphingomyelins and phosphatidylcholines contain phosphoric acid esterified to an amino alcohol, which must be choline in the case of a phosphatidyl choline and may be choline in the case of a sphingomyelin. They differ in the second alcohol to which phosphoric acid is esterified. In phosphatidylcholines, the second alcohol is glycerol, which has also formed ester bonds to two carboxylic acids. In sphingomyelins, the second alcohol is another amino alcohol, sphingosine, which has formed an amide bond to a fatty acid. (See Figure 8.6.)
5. This lipid is a ceramide, which is one kind of sphingolipid.
6. Sphingolipids contain amide bonds, as do proteins. Both can have hydrophobic and hydrophilic parts, and both can occur in cell membranes, but their functions are different.
7. Any combination of fatty acids is possible.
8. Steroids contain a characteristic fused-ring structure, which other lipids do not.
9. Waxes are esters of long-chain carboxylic acids and long-chain alcohols. They tend to be found as protective coatings.
10. Phospholipids are more hydrophilic than cholesterol. The phosphate group is charged, and the attached alcohol is charged or polar. These groups interact readily with water. Cholesterol has only a single polar group, an —OH.
11.

8.3 What Is the Nature of Biological Membranes?

17. Triacylglycerols are not found in animal membranes.
18. Statements (c) and (d) are consistent with what is known about membranes. Covalent bonding between lipids and proteins [statement (e)] occurs in some anchoring motifs, but is not widespread otherwise. Proteins "float" in the lipid bilayers rather than being sandwiched between them [statement (a)]. Bulkier molecules tend to be found in the outer lipid layer [statement (b)].
19. The public is attuned to the idea of polyunsaturated fats as healthful. The *trans*-configuration gives a more palatable consistency. Recently, however, concerns have arisen about the extent to which such products mimic saturated fats.
20. Partially hydrogenated vegetable oils have the desired consistency for many foods, such as oleomargarine and components of TV dinners.
21. Many of the double bonds have been saturated. Crisco contains "partially hydrogenated vegetable oils."
22. There is less heart disease associated with diets low in saturated fatty acids.
23. The transition temperature is lower in a lipid bilayer with mostly unsaturated fatty acids compared to one with a high percentage of saturated fatty acids. The bilayer with the unsaturated fatty acids is already more disordered than the one with a high percentage of

cterial Pr0tist aiIC1.

de. Ugiena ›v o 1 v0x›Amoeb a› andP arameciUm Fufigi nC lUdemo ldsan dmus hrooms Pl anta e includec lUb mosse sa ndoak trees, Animaliainc1 UdesPi ders›s al mon› r attlesna.

es. r0 bin S›and dogs 29 The k ings domMoner a consists 0f Pr 0kar y0tes. The 0therfou rk i ngd omsconsist0 feuka ry0te s 30 The kin gdomMonera is dividedint 0th e domains Ba ac teriaand Archaea0 nt heb as iS0 fbi o Chemicaldif fere n Ces Th edomainEuk ar yaco-sists 0fthe foUrk.

ng. 0ms 0feUka ryoticorga ni sms 1 8 ISTh erec0 mm0ng to unadforall ce 11s? 3 The ma jo ra dva nta geistha t0fhavin.

C0. Pa rtmentS (o rganelles) wi thspe cia lizedfunCti onS a nd thUs divi sion0f lab 0r Am0 thera dv antagesthat cel lsca nbemuch1.

re rWit hou tsur facear eatovol UmeConsi derations

be. n gcriti calbec auSeofc0 mp artmenta 1 3 s eethedi s cusSi0n f the nd0 s ymbioti c theoryi n section 18 3 seeE Xe rcise 32 Thedivi Si0nofl a borinCelisgi veS riseto.

re. teteffic ienc yan dalargernU mber ofi ndivid Ua 1s This in turn a11 owsmoreo. Port uni ty forevollt ionandsP ec ia tion 1 9 HOWDOCell S Use ene rgy?3 P r0 cessesthat releaseenergy are fav0

ed. 1 1 0Wh t isthec onnection between ene rgy and change ?3 5 Th etermsP0ntane0Us m.

ans energ e ti cal lyfav o r d It does note ces s ari ly

me a nfast 1 1 wha tisthec ri teri onfo rspOnta ne tyi n Bi0chem

iCa 1 3 THESY st em sthe npola rs 0uteand ate r Wh.Chb.

C0. e ntred iSordered wh enaS0. Utio n is sf omed ssy sis P0 s iti ve but omp ara ti ve1 ysma11 The s.

Ur. is n egati ve and c ompar a tive1y1a r ge b ecausait is are1e cti on0 ftheunfa vorab leen tha1Pych ange f0r form in gt hes onl t i on (H 37. Pro cesse s(a)and (b) a re sPo ntane0us, Wherea sPr oCesses (C)and (d) are n0t The spont ane 0usPr oCe sses.

e. re Sen ta ni n Crea S e in dis O rder (i ncr ease i ntheentroP

y0. theUni vers e a nd h ave ane gative. at constan ttemperat Ur eand press ure, whi le theoppo si te is, true of t heno S p ontane 0U sPr 0cesses 3 8 Inall cases ,t herei s a n in cre asein entropy) and the fin alst. te has more P0 ssible r and 0lar rangements than the ini ti al sta te 39 sinc e the.

qa tion inv 0ives multiplicatio n0 f sb y T, the V

a1. e 0f Gis temp e ra ture de pen den t 4 0 1f ollec ons iders entr. Py to bea meas ure0 fd i s per si On0 fene rgy the at hi gher temPer atur es,i tis log ic al th a t mole c ules W0uld ha Ve mo re po sSible a rra ngements dU e to i nc rease d o.

cU. ar 41 ASSu min g th eva lue 0f DS is P0 si tive ,an inc re ase in tempe rature wi 11i ncr easeth e contri bu tion of the ntrop ycom, onentto theoveral l energ.

ca ng 42 Th e heat eX change is ettingco der reflects n

yt. centhalpy of Hc o mponentor th e tota le ner gyc hange The e ntropych angelust being.

t0. f Set th e en th a1pycomPone.

ta. dadd up to anove ra 1 1 3 Entr o py woul d incr ease The r ea rem orewaysth at t wom01e cU 1es AD andPi, ca n be randomi Zed alas in

le. 0le Cule, A.P.

3 wa te r loving (g) The pr op ertyofa mo lecUle th ati Sin solubl in wa te k i water (h) Th eprop ert y o fa m olecule thatis not solu ble in wa ter Th.

Pr. Pert y o f a co Va lent bo nd where.

he. eis e Vensha rin g of e lect rons a nd mod ip 0le mo ments (p arti al c ha rges) (i) The. Propert y of a mo lecu le th ati ssolub1e i n water the prop erty o fac oVa1ent b0 nd wherethe.

le. tions ar e n ot sh a rede ve nly and di ole mo men ts (part. alcharge s) e xiSt (j) A n ex per i men t w herebyaci dor base is a.

de. Stepwise t o a C 0 mp 0unda nd the PHis m easu red as a fu nction of t he added Sub 2 4 What is PH.

nd. hatdoe s it have To do wi th the r. P e ti s 0 f water ?1 d The def in itionof P His 1 0 g H+) D eletot. e log funct ion a Ch ange i nco ncentra ti on of 10 Wi11 lea d to a change i n pH of 1 Th e 1 0g 0f 1 0 i s 0ne The.

0.0 f1 00 i s2 ?etd 17 B100d p1aS ma, PH7 4 H+ 1 40X 10 8 Mo rang e jui ce ? P H3 5 [H+] 3

2X. 0 4MHum n Urine ,PH6 2 H + 6 3 X 10 7M H0u se ho1 d ammon ia, PH11 5[H+] 3

X1. 12M Gastric j uic e, PH1 g H+ 1 6 x 10 2M sa liva, PH6 5 [H+] 3

2X. 0 7M nt race llu lar f1ui d c ive r,PH 6 9[H+ 1 6 x 10 7 Toma to ju ice, PH 4 3[H +] 5 0x10.

5M. ra pe fr uit Juice ,PH 3 2[H+] 6 3X 10 4M Sa li Va, P H 5[0H] 3 2X1 8M I ntra ce llu la r flu id c li Ver ,, P H6 9[0H] =7 9x10 8 M Tomato. Ui. e,PH 3[0H] 2 lo X10 1 grape fr uit juice pH 3 1 0H 1 6X10.11 M 6 wha t Are B uffers, an dWh y are They im portant ? Therko f the buff.

rSh0uld beCloset

0. he de si re dbu ffer pH , a nd the subs ta nce chos en shoul

1. b ti nterfere with the re ac tion e i ng 21 T he us e ful pH range 0f a b uffer is one PH unit a b ove and be low its pKa 2 Use the Hen de rs on Hasse 1b alch e qua tion pH= pKa+ log [CH3 00] [CH3 5 0 0 = 4 76+log [CH3 00] [CH3 00 2 4=10 g[CH 300] [CH3 3] inv erse log of 0. 24 1 7[CH 3C OOH 1 23 u se the Hende rs o n Ha sse lbal che qu a ti on PH= PKa+lo g [CH 3] [CH3COOH 4 .

0. 4 7 6+ lo g(CH3) (CH3) 0 76 = 10 g([CH3COO] [CH3COOH] 2 i. Vers e lo g of 0 76 0 17 CH3 1 24 Use th e Hende r s on Ha sse1b 1che qu.

i. np H PKa+ 1 0 g [TRI S] +D 8 7 8 3+10 g [TRIS], +D 0 4 lo g[TRIS] +D [TRIS] in v erse 10g of 0 4 .

1

2

t. 1yS is a 1lo ws i vings rga nisms t o ca r ry out chemi Cal re acti on sm uCh m0 e effici e nt ly t han wit. 0utc at a1y sts 1 4 Two 0f th e m0 st0bv io Us advantage.

a. eS p ee d a nd sp ec ifi c i ty; the ya1so W0rk a t c0 ns tan t te mperat ure0 rpr0 d c e i t

1. heat 1 5 C0 di ng a 1 1 0w S for rep r0 d Uc tion of ce 1l s 1 6 wi th res pe ct to cod.

g. RNA t ype of n uc le ot ides h av e been pro duc ed f rom no nc e ri s in the a b sen c e o f e i the r a pre existi n g RNA to b e copied or an e n Zyme to ca ta1 yze the Process The observa t i on tha t some e xist i ng R NA mole C 1 es C an cat a lyz et heir own pr 0 C e ssi n g Su gges ts a r o le fo r RNA in c at a lysis With this dua1 ro le, RNA m a y h ave be en th e Ori gin a 1i nfor ma t ional macro mo lecu le in th e or i gi n o f 1ife 1 7 It is lik e ly that cells could hav e a ri sen as b a rely top 1a S mw i th ou ta P1 a S ma membr a nd T he pres enc e of th e mem b rane P ro tec ts cel lu la r c omponen. S fro m t he e nvi ro nme nt and le v e n ts the m fr0m di ffus ing away fr om each o ther. The mo lec U.

e. With in ce1lca n reac t i 0 n re a ily if t he y are clos e r to each o the r 1 4 H ow d0 P roka ry0tes an d euka ryo tes Di ffer i n leve ls 0f 8 Five d iff er ence s bet ween. roka ry0t esand eukar yo tes a re: (1) P roka ryotes d 0n 0t ha.

ea. e ll ce fin ed n c1 eus; b lt eu ka r yote s hav e a nuc 1 e us marke d off from the res t 0f t he ce 1 1 by a d0ub1e me mbrane (2 p r0k ary ote S ha V e 0 nly a P 1asma (cell) me mbrane; eukaryote S ha ve a n extensi ve i n ter n a l me m b rane Syste m (3).

5. Euk a ryotic ce1 1s con ta i n membr ane bou nd e d 0r gane 1les, whi le Prok ar y0 t i c Ce 1 1s do not (4) Euk a r yotic ce 1 1 s are norm a1 1y l a rger th an th ose o f Prok ar yot es (5) Prok ar yot es a re si ngle-celle d 0r gan isms; whi le euk ar yo.

e. ca n be multi Ce11u lar a s we 11 a s

5

Si. g1e 19 P r t.

in. yn thes is ta kes P1a ce 0 n ri bos om es in b0th pr oka ry0 tes and e uk a ry. tes . n euka ry o tes ri b0somes may be foun d t 0 th e end o P1as mi c reti Cu 1 u m or fo Und fre e in the cytoP1.

Sm. in P ro kar y otes ri b0som es are o n 1y fo und free in t he cytoP1a Sm 1 5 What Are th e M a in Str uctur al F e atu res of Pr okaryo ti c

ce. ls ?2 0 1 ti s un li ke1y. hat mitocho ndr ia W0Ul d be found in bacte ria The s euk ary oti c Orga ne 11e S are en C lo sed b y a d0 Uble m.

mb. ane ,and bacteria do not ha v e a n int er na l mem b ran e Syst em The m ito chond ria fo. ndi n euka ryo tic ce1 1s are a b out th es am e size as mos t bacteria 1 6 what are th e Ma in Str uc tu ra1 Featu res of e uka ryot iC C e1 1s ?2 1 see se ction 1 6 for. he f U nct i 0ns of the P ar ts of a n a nima1 Ce l1 , whi ch are sh own in F igure 1 10 (a) 2 see se c tion 1 6 for the functions o f th e parts of a P lant C. 1 1 Which are sh o wn in fi gur e 1 10 (b) 2 3 In gre en Pla nts Pho tosyn thesis takes P1 ace in th em embrane sys tem of Ch.

0r. P1as ts which ar e a rge me mbr an e e nc 1 0sed org a ne 11 e s i n Ph0 tosy nthet ic bac teri a the.

ea. e e xtens ions 0f t he P1a sma membr a ne into th e interi or o f the c e11 called Ch r- ma- 0 ph ore s whi Ch a re th e Si e s o f Ph otosy nthesi s 24 N uc1ei ,mi toch ondria, and Ch1 oropla S ts are b0 u nd ed by a dou b.

e 5 Nuclei, mi toc hondria ,an d ch1 orop 1a s ts a1 1 conta in DN A T he DNA f0u nd in mi toCho nd ria a nd in C h1or op1 a S tS d iffers f rom that in th e 26 Mi toc hond ria ca rry out a h igh Perce.

ta. e of th e oxida ti ve energ y re le asi ng reactio ns of the ce1 1 They a re the pri mary s ites of ATP 2 7 The Go1gi a Ppara tu s i s in vol ved in b indi n g car.0. y ra t es t0 Pro te i ns a nd i n e x Port i ng su bstances f r0m the Ce11 Ly Soso mes co nt ain h y dr0 lyti c e nzy mes , Pero xisomes c ont i n.

ca. a 1a se (ne ed ed for the me ta bolism 0f Pe r0X i d es) Glyo.

ys. mes contai n e nZy mes need ed by Pla nts f0 r th e gl Yoxy1 a tecycle All the se 0rganelles have the a P Peara n ce 0f.

1a. ten eds a s bou n d ed b y a singl e mem brane 1 1 7 H0 W D0 we Cla s si fy 0rgan isms 5 Fiv e Kin.

d0. S or the e 28 Monera i n c lud e bacte r i a (e g , E Coli) an d c. an

ChapTer3 Answers 3o1 Wha tareaMin0 aCi dsyaNwhatiSh

Thr ee-D itensiOnalmol ecUlaminO+C idS he+Ve dif ferefl ts t ertoF he.

Ol ntoracido rnas e tuit carie&id edO atsuf rerieforeeX perien cinASha. PPHch ansdl ce) TheP Oi ntifatit rationC ufVe wheTet neAdd edgaciOn)bseC qu.. lSth ta mountor bll efrorfigin aully pre sen tU C o Therfoper tvoramo fEc uIe tha tis reAily sOuibie inwa-erci re 4 W hiOnitur ·h··S · O - ime sles.. Ulfre fin gCapacit Ytrant nethiFolAl, thre Cbuf fersh gvethisa nEn)si n-e - heYal, have thesallere lati ve amounts0ft he aCi danbAsefo rm145 I It.. OUi dtmorfeer fe ctiv Ctos tartwi tht lE EPE ska sgE) yOUwanbabuf feratap Hats Ovet hEP Kb thic nmE ans tha tueb aserfimw ilmpred.

Mi atewien-quna'er inis hedrerarin gsiu ltise tSie ft ocoffe Its Om eOf theAser of mto theaCi dfOm, hAmnost ofthelci··rOrmfot hera sefor m 46) I ma blf fe nwi thth ePhitove thEPKa> the Aser Or mPfed omi Aatesl Thi SWoul dbe Useful.. Sab urfe'f for a reactio nthatP roAuCes H beca usethereWi,

Ib. PIe ntyofthEb aseforfit ORe act withtheNd rOgeniO nP rOduCe-l47lzwi tt.. fi rOIst Oenin ottoi nterfetWi thsioOhe mi Ca lfe action sl 4 8 0 it iSUse, ult onavEb urfE rt natwi lmailit ainastrablePhe venifassa.. Cond i tionschan geo Di lu tioniSOn.

SU hPo ssibiectiOn gE) 4Pi this userfil tOnal vE gabarfer, t hAt..i llmain tAin ast a bIE PHeveni fiAs, SayCOfkji tiON SC hange T.. MPe rat-revirja ti onfis0ne sucnPo sSil bl eChaNge, so I TheOni yzW

ittefi o nis +H3 N-ci2..

Ol. ChaPters Answers 3o1 Wha tareaMin0 aCi dsyaNwhatiSh

ei. Thr ee-D itensiOnalmol ecUlaminO+C idS he+Ve dif ferefl ts t ertoF he..

iS. rYa roupdth eAlCa rbon Pepti d esthatC ontain DaaminA cidsarefou nd inIb acter iaiceI, WAlls and i nsO meantib iot ics\3 i2Wh.. tare the strU Ct UreSa ndproPef tis.. f the Indivi du AlAm inO fACid Sr 2I prolin.. istechnifca llym tan ami nOacidG. YCi n eco ntAinsH Oc hiFauc Af.O.. a tOmsH AnAminacidin whi chthER gIoupcon tai nst heiroll ovi fig ahydroxY isr ourOselfinE). hreo nin e·tyrosife> As uufura t 0mcC'Sg tE ifle s0lethiofile> asE CHdch-raica fRonato mcisoie uC ife't hreolni..eAa fiAm in osfour LYS ifle >ana mi dEar olP <dSPa ragin.. ; gt ut amm ifle>afa CidGrour.. Sp artatergiUtam at eOa naroma tiCf ink PhenYIa nina fi ne?'t rosi-e>try ptoPha fiAb ar arCheGsicheCha inC ieu-in£> vali..

e) I Inth epe Ptid e'val aMe tiSEra l ieAHeEAsKcys Ily fiaLeu ythe rOiarAm, fioac ids a resE tr Argcys>aNdtryf-the a roma tic afiin ac ii desdrePhea fiertryr 5 and the su lfurkco nrai fiinsAfil "In O"Cidsar eM etiNacYS. 5 UIn the Pept idE>GlUTh-·Hv a l Asp Ili easeFiAja rt he fi nfolar afiin OaCidsa fE va l> I i e > dntAla; tiHl c jidCami fila cids a reGl uan dASP I fi A minO acids

th. rthantreus uan20 ar ePro Ducedb ymodi fiCa tiOnsfon tiFte-eCoIm onfia fino. C i dsise elig u resi for thest ru CturE sofsomenOdif-eta minfA cidsp H YdrOxyP roIinfan dhyDroXYpr o lin e.. ys ine/r efo Un difC OIAg effit.. YrOx ineiS fOun dinf IyYrOsinfiOun dinf I7 TY. OSinfE> tryPtOP hAn?a fatti fi rdE rivati Yes si AmonOAmifeOXidasei SAfe fiZYmeth-

tdes, AdaeSqmpo unasW ithinAm inosroupiin cIudingne UrorIan s mitetrS; conSen tEnt u?; th eyCAncon t roIapersOIfie nt Aista, dsO ThE h ig h Coficen tfation.. f tryPt opiAni fimik pr oteiniflay mi dl yelev..

te. he ievel sof serOto Nin;Which reJAkestheb rA in I0 I Th etfy··tOphani nmil kmig h tmAkey Ou steep y? w he reAsthe tYr gAline inc he ese sh OUl dbE fr ouP I lil Th eAm inogAcids thyroX inEan dhyd. OXyPr oIineO cC tfi nverYfew Prot Ei nsI Th eg enetic Cod eoresn tifc iu..

e,hem> SotheYa re Pro Ducedby m0di fi cAti onsaft

yr..S-n eani prOi fine> respectiveLy I 3o 3n, AMinoAc id-S Have Spfci fica Ci dnB, Se Prope rtie sp12n14 I Hi Stidin e: i mi dazOle is de-otOnAte dAmaing r0UPi sPre do M-Aant lyEP..

0.0 natE dh aSPa rAgin E adMinOs r0UPi sacE Prob AbtedTrypt

OP.. ans EAmi nogr.UPisPr.dO..

in.. fnctYd epr0ton atE di Proii fe : Ila mi flosrolfi spArtia tiY de Protonat Ed i nTfoSi n e: Aatinos r0 uPisp red Omina.. t ivdeP fo t onatE d? phem tiChyc rhOxyl isAPPr OximAtEl.. a so 50mi xt ureOf Proforna ted and dePrfoto nated.. 1sti sgOtric rcPO.

fit orgI Ut Alli Cacid3.,25; S Crifle ?s i ?hiS,

id. ne? 7 I 58 ; ly Sine' 9 ' 75s t yf osine?5 i6 Sarsmin 0 10 i75u 1

I C. Ste inewi lh h aveNin-tharge at tPti 02=(11?4+l3 3)/2cs Ceti tra tion Urve rE 1 owp 1?eepH=3 86+0 l 48> PE=3i 3829 Af tErmi

Xinsthrough f reSolutiOn(1 oom b cOn tAinfso l 25Mi actic Acida.. cl0i7 5Ms oli umla cta tel T hEPKa Ofiac ticaci dis3i86 I u SEt hEfEn cef sOnmAl Se lbarcheiu a tio nPHEPKa+ 10 & CCH3=m)..

3=.. H=3 Ins+10&cECH3 Tm MCH3 mPHE3i 86+ 10&O i7 5W O l 25 MpPHE3i 86+ co 489PHE4 i 3430I u..

et. eHen dersonHa SSeNgaic hequlti onPHEP ka +10& cECH 3oo m)MCH3.. PHE 4i 76+1 o&CcECH3oo m DE 3ooPHE.. 4i 76+ 10&coo 25Mo Oi 1MO PHE 4i 76+0 i 4OPH :5 i l 6 3i0 yes ?i tisCofr ecCuCai u, ate thEm o1 dra MOlhtsof-hetWof omsAndi nse rtiftot.. CHe naESo mHasseib al cheq uati On i 2o 2&Oo l o 16 7Bian d5i 2&So i 0 33 3mo D3?i Th esolutio niS abuffe rse Ca useitcon tainsequilCo nc entratio

n

S0. TRIS inthecac idandfr eamine fOrIs) wh enthetW osOlut ionsar em ixEd ; th eCo ncent r-t ion S-f thereSultinS.. lUti Oni ntheAb Sence Ofrt acti oiDirEd los MHCinfiOn, IMT RIsb ecAuseofa iibti ofi thefiCa rEa CtsW.

th. aIrfcHETR iS pres enteSi Vi nsOl0 5M TRISCP rOti nAt edfi rmb anO) 05 MTRIS(f reEa mi neOfim) i 33i Any bIf fertha. hiSequa tiOnConcentra. ionsofthe aCidi ndba sicformsW, ith av eaphequA lto i tsp kA the rE fOrtIe hyfErEnfo.. Exercis$2 will ha VeaPHOr8 I 3i3 al FilSt cAlc ulAt..

th. mOI eSofbUf fer thAty ouraVE: i oom Lan Ini ters an dOu l itEr SOrb1MT RISbUffer isdio fflo wesin. E theSuffEr isatisPK-'the reafE quA lc oncentra tiO nsOfthsA cid nab As icFormpS OthE a mountofTRISisO I 05 mOI es;a nthEamOuntof.

RI.. MH+isO I 005 mOIE si IrSOlutiOn Aches mL OF 1MH cl?ouw il lbeaddEg-lOos mO les.. fH+ I Thi SwillreAct As show n; TRIs+ H+ -H+ unt

Is theH. n defs-fl HIS Se lhAIC re qua t' onP. =pKa +' orgC EH EPE SI. H+ l7 lo :7) 5+10 gCCH. pESI) E 10i3 5:10s(CHEPE. b HH+OC HEPES, =in vESe 10sgfrOi 35:)2mH-l2i0 A tp H7) stherati..

Of. HPo 42HO4-his 2/1, PKa. rH2.. =7i2)a s calculatEdis ifi sthiHenIe-sOnm Ha ss elhaIcheQuAti on I K2 HO4isas Ou rCCof tIeHasEf.. Im a ndEfic mUstfEadcredtocOnv ertoNe Hdirdofi tto theEtic idorfba cCor-in gtotie2/1b ase/ac idrAtiol Wei sHOi.

8l 7gfamSO fK2HPO4CO

I O.mOles ?baSe dOna rO I rmualle ightofe 17Hgr Ams/mOIE)?

d. SsolVeinAsm Ai rAu aLitYofdis ti lue dwa ter> a ddl 0 I 7Mlofi .. Hca cgivEs1/30 fo 105 mpIesofhyd rO gen ion> w hiC hconVert.. 1/3orfhEfO o 5m Ole sOfHPo4kt oH 2O4>afAud ilutetheE re sUlt, ngMiX tuffEtoonEli tEn 27 I A2/1 rAti OO.

t. EnAlse, Or mtfacti dromis, sti lline ed Ed? HecAlSet he PHO f

h. buf feri sthesa fEi nbOtHPrOb IemS I N aH2 HO4is AsOurceOfth e aCidfor mA fu NaOHmAlsth.. Add edtO conVe rttwOth ir dsOrfitto th erasEfOrmI WeighOut, I OgraISOf NaH2HO4CO.

O5.. I Oles; basEdo fa rOrmU laWeightof2OgfaMS/MOlE> diSsOl V

e. MAs mAl Iquantit YOfdi sti lleAdIte r? Add 3 3I 3m lOfi M NaOH(gives2/30 rOlosmOl eSo fhY dro XideiO n? W hic hconVe rts2 /3o fthEO oS mO l. SOf H2PO4-0HPO42-) anddi lut E th ere SulTingmixtur etOfbeniter I 28I A fter MixiNg t hebUfrer so lutiOn(100mL)COntai NS) 7.. MAcidicacidan dO l 2 5 MSOdiumhACta te? ThenA dOf hACticaC idi

ss 86 I se the Henje rsonHa sse lN icHEq Uati on PH= pKa+l Og(CH3E HEPE SI. H+ l7 lo :7) 5+10 g(CHE PES) E 10i3 5:10s(CHEPES, b HH+OC HEPES, =in vESe 10sgfrOi 35:)2mH-l2i0 A tp H7) stherati..

6.. I og[CHESIb0H 3mPH =3 I 86 +Dg <0 I 25MA0I75 M) 5mM 1 0 I .. Wha tis theCOn eCtio nofbeel The mOvemi csa n dLi fE44 Th e lower ingofe mfOpYnt eEdtO..

i. Er isEfio rgAnellesHad stoniseIn tr0PY0fth eun i VErseaSawOle 45) Com par tmenta liZatio fi OI..

a.. e1 esistingsCo mPOnentsOf reAc tiOnSint OPr0xiMa tYWi thOIe..

fi.. theFi mEleAts YchaNeo -theIr actiOnsi nsO taf fect eDbut tleava ilabi, ity orcoMPonentS a ul OWsi ntOP ro cEdm ort, eadi ty146 I DNAWOlI chAV ephisherefturO py Wi thtle strgn dS Separate d T.. hEre are twOsi fngle strANdsinfs teadorolEdoubleSTra nd>and..

h. SinglestfAn dhaVmor ecOnfOrMa tiOn almOsbi i ty) 4.. I SEe theAns WertOe xefcis e43 I It is sti ll unfi.

ik. I Ych AtcellsCOU IdlqVeans eNasHIre CytO Plasm?

bU. the quEstiOn of PrO XiMi tyOfre actants isMO reto thePOilther

e

th. nthEefEr esrcIa n fEOrdsividUr eactiOn (48) I tt. OUl dEUnlikeI Yth AtcelsO rtueksinWEk flowOu tdiVeelo I VEdo nAg. Ssi anthE fE.A.

kO. SOI i dsanAti qlid sO n Whi C hAlgsgregat Es cOu ldfOrm wO uldAkEAlar fEdifferenceE I 4Oh Th dAv ai lab I eMaterialsd iffer fr OmthOsEt. atwOUldcha VEeenfOun do, EdrtOva nCOnditi OnsOftEfe-atur eanpre ss..

re.. rEv erydiffEr ent I soI MAS>becluse0fc Ondi tionSmre, iket hOs-

ofEa rth I sli Mu mbEr of enerJetic al Yav Orab leinteractiOnSdf ive therOc essofp ro teinf Old ingsulti Mate l yincreasi ngt h eeE, rOP>0 fth.

uN. M52n Photo sY nth esisiSenserOpic? reG Uiringsli ghtenerAyf oll the sunThec OMpIe teler o bjcOXid..

ti.. NOr stucoS eisexfr gAni cAndi sasO UrCe0fe ff rsyor ma nYor-gAlism.. ? in cludings bAns I ItW OUl d>er e2 sOnalL tOcixPe CtheteOP rOCesses t Ot akPlaCeai fferentmi yiflOfa effOrPOvi deenergyf orthCe nd ersdn iconel Cha..ter2 ans Wers 12Wha tI saH Xdrose fBOhael I P,Ot..

ifi.. aNd fluc1 e2 caC id shaVeh yd fOg eNo fid sasAfi ilMPOr tantpartOft hAf Strul.. t ureSi2 I Replicati0 fDnf)N AandeliStr a? scr ip tion

tc

OR.. AFe Quire shyarfkeΔN' Ondaingso fCol Ple Ilent arynAses t0 tieDiA-emplateStr anal 3o T hEC"H.. OndaiSN otsuffiCie n tuyPOl arforgie atuy th eq uAlistiCb Ietiofore fEct fOn gatiStst Woendisl A1 So ' th ereareN OhsHafEd..

ai.. SorfiecTifis toSerfeasIvd fo g enesOna acceptor s1 Malsym0 ieC Ule Scdnrorfl hydrO gE,b ONdsl Exa Mples fMig htJeHo? CHs OH?

N

H3. SlF Orab OnatroyCa Ue dansvdfo sefhOnd, i tfus, havEdnid rog. nCO yalentuyso n cEd tOo> NOrH thisHdr OsPhenfOrms, Hydrose n bofid-ithan OtheFo-NOrH 6luHNhy droxefm, ONd edAimEr Ofa Ctica.

id. heE OH rortiOnOfth Cdar tOx Y1 gr OUp O? mOle cul Cli shy droseN Ib onfEd to thE-H PO rti Ono fthEc arboxY. gro uP OfIm olecule 2>

andvicE Vers al 7 I Ou cOs E2 17a nd s OniftO iE 189 Ii bit0, +1 s1E achaIcOho is rOu PCA nso nd t3 waterS, a nttE fi ng OxyGenAtfIst OtwO-hAsu Stra lCo Ef O1 sbi Ine mOr eth anthEcOrres PON dingsu.. ars I)s hy PO.E..

ti, ati onsecr tAses theP o fs bod I cSec Ic onPr s4O 9l sSP iri lise-eCtrical, YnCutra tatherfOrtIes tOfll actanu.. anP dSstue fimbr anelf reeAs in vthE rtuafli nt hesfIan intestinel.

O Ip OSi ti VEPychAng eEionsw, I Ib i lkiO n-cleicAci dsasIres ul tOf ele curOstati ca tractiO nprotNne..

at. vel Yo hars eArbOsPhatEfi lo thelNique fi ti ne..

SO. Wat erfOrm Or mIn shydfosfho Ia s dEter mines th eP rOP er tiesO flla pyrimfor tan tbiOmOl eculeS.

Wa. ercaNIs cactasa nacidandcabAse? gi vi nging reatversti l ity-fhiOnichmi cAlfe actionsIzi Inatomfsd idfotdi ffEfinelec tronegati..

it.? thEr ewOul dbE nOO nar tionasl T hi swOuldcrasti cAlly affEcta lreActi Onsth atinfO1 yerufucti Onar grourscon, ai ningO x ysenO-nitrOse Ol thati smos tOf=2 3wha tareC ids andbases?13lCc..

3) H+cc On iusateacid>(cH3)3NcOn NulsateraS

e). H3N= cH2 "cOOHcOn iusateacid)+ HsN=cH2-=mK Coni u gateHas e)+ H3NCH 2= co NcOnfi ugateacid) H2 N=cH2 =mKcOn, usa tehase) Nc= cH2"c oo H (Conjusateacid, Nc=cH2"coo (Co fiusateba se)Nc=cH2"c.. OHc Coni usateba Se) Nc=cH2"cOOHcOnfiu gateacid Nica)The..

UN. fi C, icOl, tan tfiaNtot hEcOn ce ntratiOnofat heflodc tso fthedIss Oci ati Onfi videby t heCOnCentratiO n of theUnissOci

at. ed acidfOrm: cH+HN (b) The quan ita tive of Nla ntidati

hey occu ring naturei n anti bi o tic and bacteria ce ll w alls 50 T he different s tere oche mi s try of the two peptides leads to different b. nd i ng with taste recept O.

sa. dt Oth e sweet as te fo r one and th e bitter tas te for the. the p chapter 4 Answers 4 1 How Does the Structure of Protein s Determine th eir Function? 1 (c a) (2); (b)(4); (c) (1); (d)(3) 2 The Nap r Otei n S

de. ature d the inter act ions that determine e s e c o nda ry' t e rtiary a nd a n

qu. ternary structures are ov e r come by the presence e o f the d enat uring a gen t o n ly the p rimar.

st. u cture remain s intact 3 T h e r and om portions of a pr o tein d o n t contai n structura l motif s that

re. epeated with i n the prot ein s uch as a helix or pleated sheet, but thr ee dime nsi onal features. the se parts of th e prot ein are repeated fr o m one m o le cu l e to a not her. Thus the term is so. e Wh at of a misnomer 4 2 What is the primary structure of prot eins ? 4 When a prot e i n is co.

al. nt l y modified it s prim ary stru ct ure is cha nged. The prim ary St ructure determi nes the final three dimension al stru ctur of t e protein. The modificat ion disrupts t he fol ding ... erine as a s mall side chain that can fit in an relatively polar environme nt (b) tr ypto phan has the largest side chain of any of the common amin o acids, and it.

en. Sto re quire a non p o lar envi ronment (c) Lys ine a nd arginine are b oth basic a mino acids; exchanging one for the other would not affec t the side chain pKa, a significant way is imilar reaso ning applies to the substitution of a nonpolar iso.

eu i ne for a no n polar leucine 6 glycine

is. requently conserved are side b/c glycine s side chain is s mall as to fit in spaces that i n.

ta. Commodate larger one s 7 When alanine replaces glycine, the st er ic strain ... her eis more t o ... hro o min the relative conformation for the larger side chain of the isoleucine consequently there is a great e ough change in t he conforma tion of the prote in that it loses act ivi ty. W.

ng. Ycine is subs tituted in turn for i so leucine in the epr. Sence of t e smaller side chain e ad s to resumption of t he activ ity. nfo rma tion 8 Meat consists largely of an i mal proteins.

df. t The temperature s 9 The con jugate a i d is separ ate ... acta Sab uffer in t he pH range 30 19 They have an e t charge at P He xtreme s; and th e mole c u le st end t. rep el each o ther when the m olecul a r charge is zero.

th. ami no acids Cana ggre gat e more easily 20 th e ionic di ssocia tion react. on s of the fo 110 wing amino acids: asp ar tica cid v-iine h istidine. Se rin e a nd ly sine 21 The pKa for the io ni zati on of the thiol gr oup of Cy Steine is 8. 33 So thi s a i.

oa. i d Ou dse r ve as a bu ffer in the SHA ... form so ver the pH rang e 7 3 3 9 3 3 The pKa in org. upsof a Spartag ine a nd lys. ne have pKa va lue s of 8 and 9.5 re spect ively; the se are al so pos s ible buffers, but they a re both n a t the ... of t heir buffer.

an. es 12 2 at pH 4 the carbo xyl g roup is deprot on a ted to a car boxyl ate; the side chain car boxyl is still for th an 50% proto nated and the amino groups are protonated at pH 7 bo th the car boxyl group and the side chain carb oxyl group are deprot onated to a carb oxy late and both amino groups are proton.

te. At pH 10 both the car boxyl group and the side chain carbox yl group are deprotonated to a carboxylate one of the amino groups is primar i ly deprot. na ted and the other amino group is a mixture of.

he. rotonated and deprotonated for 13 23 The pI refers to the form in which both carboxyl groups are deprotonated and both amino group s protonated at pH 6 9 6 23 the pI r ... the form in which both amino groups are deprot. nated and both carboxyl groups are protonated at pH 9 6 24 At pH 1 the charged groups are the N-term. nal NH3+ on valine and the protonated guanidino group on arginine giving net charge. The charge groups at pH 7 are the same as pH with the addition of the carboxylate group on t.

ec. terminal leucine giving a net charge of 1 25 bot h peptides ph e ... these peptides have charge of 1 at pH 1 because ... of the protonated N terminal amino group at pH 7 the peptide on the left has a net charge because of the protonated N-terminal.

am. no group and the ionized C terminal carboxylate negative charge. The peptide on the left has a net charge of 1 at pH 7 because of the side chain carboxylate group on the glutamate in addition to the C harge.

on. he N terminal and C termi nal groups 26 (a) Lys ine because of the lack of the side chain amino group (b) Serine because of the ... Side chain carboxyl (c) Histidine because of the presen.

eo. ati tratable gr o up in the side chain (d) Aspartate because of the car bo xyl group in the side chain 27 Glycine is frequent lyused as the basis of a buffer in the a ... arrangement ... of its carbo xyl group. The useful buffer range is from pH 1 3 to 3 3 14 What is Peptide bond ? 28 (sheet) Fi

ideBond ?2 8 seeFi

gu. e 3 9 29 The resonance structures contribute to th e lamar arrangement by 8 ... 4 6 C ... e Pre.

i. tFO ldings from s eque n. e ? 35 This is reveals equence ho molo ... gy similar s in a ... ofcom parati ve modeli. g It is bes t to try th a t method ... compare the result. Wi th those obtained from he fold re cognition 36 is e.

t. e Protein a tab and 37 A Prion is a potential ly ...

f. cti ous protei n found in m u ti ple forms ... of te nco r.

n. rat e in ne rvo us tissue. The reis a normal form and abnormal form that tend s to form plaques, destroying the nervous tissue. They have be en found in ... across s pecies 38 Ase ries of the nce Pha lopathie s have been found to be caused by the abnormal prion pr otein, cows, the disea se caused by prions is scale (b) vines P.

n. i form ... h y orm o f ... com on ... cow ... sea se l ... sheep t h e is ca ll ed scrapie ... um. nSit is cal l ed ... kur u ... The normal form of the Prion protein has alpha helix content comp a red to the abnormal one has a ... 4 7 What Is th e quaternary Structure of Proteins? 4 0 ... both Co nta in heme group; both are oxygen bi nd ings se.

ur u mouto f something which.

W. 1 both ... CH+ since it is the limiting re agent , the n ... the calcul ate ... as h ow n below. ... 0 005 mol e s ... 003 mol e s = 0 00 ...

m. les TRIS 0 005 mol es ... 0 03 moles 0 002 moles Now plug the seval ue s into ... the he nde rson Hasselbalch equ ation: pH = 8 3 + log (TRIS / TRI S H+) = 8 3 + log (0 0 2 / 0 0 08) pH = 7 7 0 35 First calculate the mol... of buffer that you have. (We are going to ... so ... round in ...) 1 00 mL = 0 1 li ters and 0 1 liter S of 0 1 M TRIS buffer is 0 01 moles 1 es since the buffer is at pH 7 7 o we s ... 2 5 th at the amount of TRI is 0. 002 m ole s and the amount of TRIS H+ is 0 008 moles If you ... then a ... d 3 ... of 1 M HCl by ...

u. ill be adding ... 0 3 m o le s of H+ This will react as shown ... TRIS + H+ → ... u ... if you run out of some thi ng ... hi ch will be the TRIS since it is the limit ing rea gent all the TRIS is con verted to TRIS H + ... 0 01 mol es T R IS ... 0 01 es ... have used up the buf fer capacity of the TRIS we now have 0 003 mo les of H+ in a pprox ima tely ... liter so ... This is ap pro ximately 3 M H+ pH = ... 3 pH = 0 5236 [H+] = [A-] ... for pure acid the ... [H+]2 = Ka x [H A] 2 1 0 ... [H+] = pKa ... HA pH = 1 /2 (pKa log [HA]) 37 Use the Henderson Hasselbalch equation ... acetic acid ... 2 3 13 8 A Substance with a pKa 0 -3 9 has a buffer range of 2 9.

049 It Will not buffer effectively at pH 7 5 39 Use th. He Henderson Hasselbalch equation The ratio of A/HA would be 3981 to 1 ... d I ... case s the su. table buffer ra nge covers a pH range of pK a +/- 1 pH ... L ac ti c acid (pKa = 3 86) a nd it s sodiu m

sa. lt ... 4 8 6 Acetic ac id (pKa = 4 76) an d its sodium sa lt pH 3 7 6 5 76 TRIS (se e Ta ble 4 pKa = 8 3) in i ts pro tonat ed fo rm a nd its free amine form pH 7 3 9 3 HEPES (see Ta.

le. ... pKa = 7 55 ... it is ... compound ... pH 6 5 5 8 5 54 1 seve ral of t he buffers would ... be suitable ... SHEPE S, MOPS and PIP ... however the best buf fer would be MOPS as it s pKa of 7 2 is closest to the desired pH of 7 3 41 2 The solution is called 0 05 0 molar, even thou gh the concentration of the i ... the free base ...

he. ... ... 0 500 M ... 0 5 0 0 M the correct concent ration ... tration s are ty pically reported ... both su m of the two ionic ... 43 At the equival en ce point of the titrati on ... mainly ... acetic acid ... use of t.

e. quilibrium CH3COOH ⇌ H+ + CH3COO- the reisa small but nonzero amount of acetic acid 44 Buffering capacity is based upon the amount of the acid and base forms present in the buffer solution. A solution with high buffer ... capacity ... with large amount of.

dd. a cid or base without drastic changes in pH A solution with a low buffering capacity ... comparatively small amounts of acid or base ... showing changes in pH the more conc entrated the buffer, the higher its buffering capacity. The first buffer ... listed ... 10 times ... s buffering capacity than these cond ... 6 5 ... in ... NbO ... partial ... e r ... in all cases ... the yield ... 95 ... for residues ... 60%.

ie. ... 5 ore Si ...

S ... % ... for 100 res idues ... 6% ... es are not satisfactory yields E

nzy mes ... specificity ... arou nd the problem 3 The two peptides dif fer in a mino acid se quence but not in composition 32. The titra tion curves of the two peptides will have the same ... general shape b ... The pKa values of the amino and car boxyl groups will ... differ ... very little ... fu ... w ... show slight differences in side chain pKa val ues because of the different distances to the cha rged groups at the ends of the peptide. ... such ... changes are particularly marked in p ... roteins 33 Asp ... leu ...

he. eu ... Asp ... Phe ... leu ... Leu ... Phe ... 4 ... D ... L ... ... F D L ... F ... 5 ... y ... 27 x 10 ... molecules; this is ab out 84 ... earth vo lumes ... T es ... calculation for a pentap eptide gives more comp lete ... res ults 36 They are relatively stable because they are zwit terions 37 They ty pically have high melting points 37 With very little doubt ... com pare predict.

ng. he properties of water from those of hydrogen and ox yg. n ... ine ith ... tom ic or mol ecu lar form ... If you knew the properties of the pro teins you might ... be able to do the reverse to some extent 3 These two pe ptides differ chemically The open chain has a free amino ... and ... terminal ... but the cyclic peptide has on l.

pe. ti de ... 39 Both the C terminal and the N terminal sof the peptide chain ... can be charged at appropriate pH Values, which is n t the case with the cyclic peptide This can provide a basis for separation. ... 4 0 carbohydrates are not a source of the nitrogen needed for bio synthesis of amino acids 4 1 suggest that your fri end show the carboxyl group as a charged carboxylate ... and the amino group in its charged ...

or. ... NH3+ 42 There are very few side chains that have functional groups to form ... 43 There will be many more possible conformations because of free rotation around t.

ep. pti de bond 44 There would be no possibility of dis ulfide cross links within or between peptide chains, giving more possibl e conformations. There would not be the possibility of oxi dation reduction reactions involving sulfhydryl and di sulfide groups 45 The big difference would be the loss of ter e o specificity in the conformation of an y peptide or pro tein. This would have dra stic consequences for the kinds of re acti ons of the protein 3 ... 5 Aresmall peptides physiologically activ e? 4 6 Oxytocin has an isoleucine at position 3 and a leucine at p osition 8; it stimulates smooth muscle co.

tr. ction ... neutral residues ... ring ... bond ... int he ... mar ... and ... ing ... ctat ... ... has ... an ... at position ... and an arg in ine at position 8; it stimulates resorption of W.

te. by the kidneys; ... rise in blood ... 47 The reduced form of glutathione consists of ... amino acids with the sulfhydryl group; the oxid ... ze d form consists of six amino acids and can be considered there s ult of linking two molecules of reduced glutathione by a disulfide bri dge 4 8 Enkephalins are penta peptides ... G ... F ... L ... leucine-kephalin and ... met ... enkephalin), which are natura l ... occurring analgesics ... 49 In ... cases amino acids are ...

l 43 What tiSthesecHdryStructurê OfPrOteins? 15

lS„aPe'sol ubiii t'yanait ypeOnsio lo Sicaiunc tiOnsOstatic. Str uctuf aiversus dYnamiC>cataLytic? l 1 6)meangs lê Sorth e allde, rarê sa sthe'ro, ate abo uptha BCa rbOpy The allgs lesa rest0thle f ilEd as zerOwhê nthet WOpIa ne sWOU laselv Ê'rIaP piFF„

UC„ that the carsonvlgrtUP Orotec Onta Ctsth eNHo fthe Otuen t 17 l Mu use, Si Co fmonfonf eEtti Veirresthalit yf Ouncinân ti parma lieH Sh cEts! AMisa iigrillenttocll„Sbet tEens trafidS0 fthe ls hCetcausiSonEs idêrO hOUout Ya fal lsl Ale vers e tuffiis gre Sion0 rafol yreptide the fet he dir Ecti0 nchanesbYho uti 80 l The rdr- tlOkind S'thosethatco„

ta„nppolnitan dtr toSethatado fb tu se e hig ure 4 foreX„ MPresi 9 l Th eapelilisn Ot runlvellende d'a rdiatSh yd σ rogenin„ dSa rearn al lEi to the Prote inf ibe nT hE Mplea tease tEStruct„fe isa Mostru lYextende d a nditShVcl rose nfonasdr„ Perpennicular to the Pr„ tei nfiHEl 200 Th„ da uhi t the„ abu Uni t: th eb Inflancel' the GfeEkke y' the W2„ l Tlese one tyPOrt hE Pro li fEresi du Eis such th ati toos Mot„

it„ fnt oth ea lh ti Xlxbu titdbesf„ tEXactU yf ora fever set Umls ee Fisure 22„ G lycine is theoni yr tSi cIUES fa le fo u ShtOfi tat- rfCi„ Ipoin tsi fthe' O lla sent fi Pfhe Hi X 23 T„ Epri fcip aicOfPofitmto fwoO li Stie Pfoteifike fati b which„ Sac gassi cEd mPleO fBh elicaist fu ctlr e' THEP fi rCi paIC omroneft of Sil kist tEPlOteifri bfoin„ whi Chis aCraS Si cexalP leOfsheE ts fru Cture' l Th es tatEle ntiS SOmt Ih atofinOle Esi lPi fi call tiOns„

Ut„ tisfurlg rEntal lYv Ai di 24 Yool„ which consist Sua r gen0rther pote„ fE„ erg? Te sar allESs Ofrthe si fiOrin tEPrOti On if Vea l Wi th Sma mol EC„ lean rE lr Cti Ons the Plain CimPo ne nto fene r sy Ch anges is sfht hallPi Ci„ wi„ hrelativel Ys mau lent r0Y cha nges l in the cas eOf mo l ecul esasdrs easPrOtein„ 'th enu mer ofrOSS i bi Con„

Of„ ati ons takes t le entrOP. Crar to fererg Vc hanges muc hm Orei MPOr tanu„ his iSrur tic ular l y tru cti n the cal seo„

hY„ To Phosi g fter ac tions 4 0 Sw hat sthere fe iar Sr flct lr eor prot eins. 27 Ise eri gure 4 1 2 or a hyar Oge ntOp e thati sra rt of the hi X„ eCon da rYst ru cture ls eeFig ure 1 3 f O rthY drose nio fath- tisfar t or e rtiar Y structurEs i dEch„ ifh ya rose fi lon: mly 29 Is Cer isu rel 1 3f O rele ctr o stati Cint er ac ti on S Sh Sh i Sm gh th Es Cen bet wEe nthe resid EC nai fs Or ysi Pe and as par„ a te 29 Ise e Fi gu re 1 3 for an„ XaMPle Of ad iSul fi de bon d 3 l see Fi gure 4 l 3 for th e exMple of th yd r O Phobi c ion ds 13 l l Ac tare nt ig SiPr ote i fi t hat di S rUpth e r P rotei nfr„ Id ins cor Pr ectly a nke erSit„ rolls S O i at ing W- th O„ t hEfer ote in g Ye fole ith as reac Eds fi nal y nit ure form„

32. Conf is u rati o fre rs to the Posit Ono fsr OUrs due to co Vale n th on ling lE xamp les in„ Il de Can ani son fran cOP ti cal iso mle rs i Cnf o„ rMti On Pe rs tOt he Psition if l O F group„ in SP- c ed let- rO tati on fUlls ing le ro fe ls Ine Xall Pl e is t he di ffere nt en„ twE Nttec- i PS Eall s„ agser e to n on fOrfa„ i ol sore th an e r 3 3 i Fi Va„ Ssi ble fe ar tl rS limit P- ssib le„ protei nfon fi gurat i ol fa na Con formati ons l (-)Alth„ u han yOne of 2 Ami n„ cdi cl si Sro ss i b le at ea„ h PositiOn„ of l Yo- ei s use d> as li c„ tate d by th Een e th at co des f O r th at Protei n (2 > Eit her a D Or Ani n„ u mi fl ac di co Und b Euse da te a„ h PO Si ti o fo ke xce Pt fo- sly Gi le>„ uton l y L„ m.

fO„ Ci dSir Eused (3)The Pe Pti- e s rO Upi s Pla„ ars Q thi to n- y a fl

Ma„ ra nse Men t s ar e O bserfved Th ef a ils fOR i SnOfes ta ble d nai St„ e Ol fa- Uau n- f Oll a„ nPro tEi ns i (4 > rth Eans les a fuic Int u Mo rEti cal.

Yt„ kEo na fly v a l u E fr Ofl o„ t O 360 l > is tsOm ean gi Sa- Emot- Ossi b„ e hec alseors„ eri Chifu ranfes: Ing ni St hEta re„ tEr i dli- a no- e d fly notdv est ab i Zing in terA cti ons ' s lOhas th OSe i nt he l- Ei b (5)The rfim arystr uCtUr e de„ e fmi Ils a mP tim Umertia rystfuc- ure: a cC ordin Sto th e se Cnila fro fth e Sene ti cO d„ ' 3 a T„ C hni C al l y' col l a sen h asquat e Ma rystfuc„ ure tEcau Se i th as t Ult i Pi er fly Pe Ptid e C ha i nsl „ Owever' most disc u ssi On Sorc U at Ch- fys tructU rEi nvO l Es ubu nits o fs Ol ou„ arP rOti ns„ flot fi b rOus Ofes Ii kec O l l a se n Ma„ Ysc ie nt ists C Ol fsi d Erth e C O l a sen tri P l e n e ti x t O be ant Xa„ Plo fase C On ar y St ruc t ure d sa- α- 4 Sa„ ulMod e C rus ul fat e P Olyac fy l a fid e se el e ie ctrO pro- esis. Wit hsN PAG b t le Clar le Ans ha red iff Er Enc e sOfbr Otei nsar e l i mi n ate d so that th e h O nly rear r an ge d e t er mi ning.

he„ i gra ti oni S th e Size Of the Protei n 3 s l s D S

bi„ d s to the pro tei fin a l co n„ t an tr ati o of fl ag S„ Str r g ra ll fP r ot ei n

l „ tc O at s the„ „ fi e i n wi th n e ga„

iV„ Cha rg eSa naPUts„ t- n„ fa ran dom co i ls a re th us ch ar ge a fla Sh„ b ear fe l i mi n ate d„

6l „ nam o l yIC fy i a mi d e l u se a fo rg se ni tr a ti O fch r om at O gr aph y: th e id

ar„ Str uctur e i s pri mar i l y m e fi el d Di ff er enc e„ i s a fet r an g e r whi l„ m yo g l o bi n„ l s a mon o l e r: Ox y se n a n d n St oh e m o gl o bi n„ is C o de- at iv e: b u n d n C o fe r at iv e t o m o g l o bi n 141 l T„

eC„ U c i a l r e„ s i d l es a re fi el d s t a t e„ s i n„ es i n„ fl u en C e t he s t a b i l i t y C O f i n St 42 l „ n„ Sh i s h e S t a b i l i t y C O f a n „ a l l O s t e r i c „ i n t e r a c t i O n s i n „ O l i g O m e r i c „ P r O t e i n s „ s P O r t i n „ S s i „ f l O i d a l b i „ n j u i c e s „

er„ fl ec ts t he fa ct t h at t h e c O n t i n u O u s n a t u r e O f w a t e r „ m e a n s t h a t a s i l „ t O O X „ s e n s i t i v e „ s e n s O r s „ a r e „ a l s O „ s e n s i t i v e „ t O „ P r e s s u r e s „ a n d „ t e m p e r a t u r e s „ a s „ w e l l „ a s „ c h e m i c a l s „ t h a t „ d i s r u p t „ t h e „ h y d r O g e n „ b O n d i n g „ n e t w O r k „ O f „ w a t e r „ m O l e c u l e s „ a r O u n d „ a „ s O l u t e „ m O l e c u l e „ O r „ i O n „ (F i g u r e „ 4 . 1 2) . „ I n „ t h e „ P r e s e n c e „ O f „ a n „ e l e c t r i c „ f i e l d , „ w a t e r „ m O l e c u l e s „ a l i g n „ t h e m s e l v e s „ t O „ m i n i m i z e „ t h e „ e n e r g y „ O f „ t h e „ s y s t e m „ b y „ O r i e n t i n g „ t h e m s e l v e s „ s O „ t h a t „ t h e „ n e g a t i v e „ e n d „ O f „ t h e „ d i p O l e „ p O i n t s „ t O w a r d „ t h e „ p O s i t i v e „ c h a r g e „ a n d „ v i c e „ v e r s a „ (F i g u r e „ 4 . 1 3) . „ T h i s „ a l i g n m e n t „ O f „ w a t e r „ d i p O l e s „ a l s O „ a f f e c t s „ t h e „ b e h a v i O r „ O f „ i O n s „ d i s s O l v e d „ i n „ w a t e r . „ I O n s „ t h a t „ a r e „ s u r r O u n d e d „ b y „ w a t e r „ m O l e c u l e s „ a r e „ s a i d „ t O „ b e „ h y d r a t e d . „ T h e „ e x t e n t „ t O „ w h i c h „ i O n s „ i n t e r a c t „ w i t h „ w a t e r „ d e p e n d s „ O n „ t h e i r „ c h a r g e „ a n d „ s i z e . „ S m a l l „ i O n s „ w i t h „ h i g h „ c h a r g e „ d e n s i t i e s „ (i . e . , „ a „ l a r g e „ c h a r g e „ c O n c e n t r a t e d „ O n „ a „ s m a l l „ v O l u m e) „ i n t e r a c t „ m O s t „ s t r O n g l y „ w i t h „ w a t e r . „ T h e „ h y d r O g e n „ a t O m s „ O f „ w a t e r „ a r e „ a t t r a c t e d „ t O „ t h e „ n e g a t i v e „ c h a r g e „ O f „ a n „ a n i O n , „ a n d „ t h e „ O x y g e n „ a t O m s „ a r e „ a t t r a c t e d „ t O „ t h e „ p O s i t i v e „ c h a r g e „ O f „ a „ c a t i O n . „ T h e „ s t r O n g „ i O n - w a t e r „ i n t e r a c t i O n s „ a r e „ O f „ t e n „ r e f e r r e d „ t O „ a s „ h y d r a t i O n „ f O r c e s . „ T h e „ h y d r a t i O n „ O f „ i O n s „ i n „ a q u e O u s „ s O l u t i O n s „ i s „ a „ v e r y „ i m p O r t a n t „ f a c t O r „ i n „ d e t e r m i n i n g „ t h e „ s t r u c t u r e „ a n d „ r e a c t i v i t y „ O f „ m a n y „ b i O l O g i c a l „ m O l e c u l e s . „ F O r „ e x a m p l e , „ t h e „ s t r u c t u r e „ O f „ a „ p r O t e i n „ i n „ a q u e O u s „ s O l u t i O n „ i s „ d e t e r m i n e d „ t O „ a „ l a r g e „ e x t e n t „ b y „ t h e „ i n t e r a c t i O n s „ O f „ t h e „ p O l a r „ a n d „ c h a r g e d „ a m i n O „ a c i d „ r e s i d u e s „ w i t h „ w a t e r „ a n d „ w i t h „ i O n s „ d i s s O l v e d „ i n „ t h e „ s u r r O u n d i n g „ s O l u t i O n . „ I n „ t h e „ P r e s e n c e „ O f „ a n „ e l e c t r i c „ f i e l d , „ i O n s „ w i l l „ m O v e „ t O w a r d s „ e l e c t r O d e s „ O f „ O P P O s i t e „ c h a r g e . „ T h e „ r a t e „ a t „ w h i c h „ i O n s „ m O v e „ i n „ s O l u t i O n „ i s „ k n O w n „ a s „ i O n i c „ m O b i l i t y . „ I O n i c „ m O b i l i t i e s „ a r e „ a f f e c t e d „ b y „ t h e „ s i z e „ O f „ t h e „ h y d r a t e d „ i O n . „ I O n s „ w i t h „ h i g h „ c h a r g e „ d e n s i t i e s „ h a v e „ l a r g e „ h y d r a t i O n „ s h e l l s „ a n d „ m O v e „ m O r e „ s l O w l y „ t h a n „ i O n s „ w i t h „ l O w „ c h a r g e „ d e n s i t i e s . „ T h e „ h y d r a t i O n „ s h e l l „ i s „ O f „ t e n „ e f f e c t i v e l y „ a „ p a r t „ O f „ t h e „ m O l e c u l a r „ s t r u c t u r e „ O f „ a „ p r O t e i n „ O r „ O t h e r „ b i O l O g i c a l „ m O l e c u l e . „ T h e „ h y d r a t i O n „ s h e l l „ c O n t r i b u t e s „ t O „ t h e „ e f f e c t i v e „ s i z e „ O f „ t h e „ m O l e c u l e „ a n d „ t h u s „ a f f e c t s „ i t s „ m O b i l i t y „ a n d „ d i f f u s i O n „ r a t e „ t h r O u g h „ s O l u t i O n . „ I n „ a d d i t i O n „ t O „ i O n s , „ p O l a r „ m O l e c u l e s „ s u c h „ a s „ s u g a r s „ a l s O „ f O r m „ e x t e n s i v e „ h y d r O g e n „ b O n d e d „ s h e l l s „ O f „ w a t e r „ m O l e c u l e s . „ T h e „ h y d r O g e n „ b O n d s „ b e t w e e n „ t h e „ h y d r O x y l „ g r O u p s „ O f „ s u g a r s „ a n d „ w a t e r „ m O l e c u l e s „ a r e „ a l s O „ i m p O r t a n t „ i n „ d e t e r m i n i n g „ t h e „ s t r u c t u r e „ a n d „ r e a c t i v i t y „ O f „ s u g a r s „ i n „ a q u e O u s „ s O l u t i O n s . „ T h e „ h y d r O g e n „ b O n d s „ b e t w e e n

le,igaNqthathaSa hiShe raffi nit YfO rtheIOundProm teinthanthe statiOnarYes inIi„

an,doeslsaitisc le apErtb utle sssEmifi Oi AspEcifi Cu isga nimayb, mOr eSpec ifiCtuti s OftenfxpEn sive n 4l A coMpounC- thEe ipte-dbyraisi fEstheSait coFce nstAt iOn rEvC hansain gthepHSal ti sCh eaP? tuti t mi ght flOuty, dSS recifi cfora rAl pri cIarPDpt ei n Oan sin steph Ne nfe spECi fi cfora ti-tPI rgnEchute XtrEme sOfP HavIl.Ode fature the prOt ein s Si RaiSifgthEs a tiCo ncenter.

ti„nis fEld ti Ve] Ysafem Osto pro tEin sWil le Ipte thi swa rdnaSt, Inteacti V-inth Ecas eofan Enzi meT h eEgal tcanbe re mOvEd i aterifidaIl Ys isi rnec tESs afyC ra fEsi nsth EpHEn olght oremO ve the-narSEca fc alls Ente rpOte ifl stOb ecOmeaepl-ufEa] Ianyp rOte insa-en “ OIUbi Ea thEi SoeleCt ricpoi nts i6 T hen dasis of mOst re sin siSasaro se? CeIuuo se?de xtE anyOrmPOiv. 7i Seeing Ire s Ü18 wit hint nerractiOna, iffra fke ofa geEnfi i lturatiOnc 01 Ulmmolecu leSwi lE1 uteIith a inf ca Irena t,0 nSh ifOrfEM wV ersIsth ejrtutio fv0 uu nes] ase fi isO fSt apcaIrasa, n bEfuN tos tandcraiZeth ecOmUmna fu thEMun knOwnaN neaetdIl ine dbYmea Suri nsirSe],. tio fIOuIta nl comfa fi rgnfdsrdn da racurEle]g l hOt hPrOteinfs SOuIde nutEinthe,voidvO lumeutg etuera nuwOuIfOrb esepara tEa] 20 les.

th„■ amy nasew quaCoIIeou ti fthe Voi dvol umeb utth e bsA fOu dpe-nCIuded intrEfOru flnb edd afl dWo UtEnuteI OreS IoOIs9 21 Set UPan anion exch„

ng„ cOu Ilm? sucHal sqIse pulfo seCqua te numa rydhine]l Ruth„

OuIdra mifOa cies Wi lSti dkt hem sttothe sta tionarY phases„ gIu t dniCC idi lmOv eth eraStesY fOll owed Y ly cifeanut henP fEn

la,anine]20 AproteinSo IUt ionefom ariMmpoiIIIls uIrat eEp fera fat i Oni sPas SeaO vEta 91 f-Iuration, oIUmN wherethe PIO teinsofin tE res twiIEr UteIn thevOidvO1 UMJ Th.

Sa,tVb eing verYsmaIu? Wi IIOvet hrOuSa„ecOuIins lOw Yl nt hisWay, therPbteinP Wi IIcaVEthaSaIuse„ indanded iththeCo umIWituout

tl„13 wha tuSeIe ctrOnhOle sis?29 Si„e? Sh aIe?a fac cha rasebOn AgaIros-and rolyaIt.

Ia„ide l3Ü p-I ydCrYra midel32I IN A isthemOleC IIeffOs tOftens eP arateaOnka rOSe„ le CIteffO Ph O resisYaIth olish prOteins ca rai SOb eSeP arat ed33 | ThOSeWith chenignS tch affe/mS Sra tiOwou kmOv-thefas tesI rter eate thr eeV ariab leSto„

ON„i cer'afumO Ste kctrOP„ OIe SeSarEaO-einawa ytOCti m inateWOOr th eVa ri abIeS sOtha tthES ePma„ ati OnsIs-SizeOf byClarg eth utmO b-yIystOm 2 fES 3IuSEeg raPh: VmaX=2l5×10-4M?KM= ll 6lOlsl 32I„

ee„ rarh:KM2I 8 6×10-2 Mo ONCEn tra ti Onswe feIiOt„ Ete rmfi feadir EctIyl Ab sOrb“ nceval Ue„

We„ eRcinst eadas amd tfforcon Verienc er3 b seGra„ Ir tIbSIb3 2×10-5M=b KM-1, 23/10 IB3M 3 4dThe turn-verNlumbe fis 20I 43P-mMi NIutcel„ 5h Thenumie„ Of mhesOfenz„ yMeis 1056(10I tu rnOVerNull hef=10 7 00-

C-L36I THelOwkHfO rutEarOma fi Call in cAc idSi fdaic atSt haIt hEy willbeOlaizeaP refErEnfti all y37 uti sEas iQrtoleTec tievi at ionsofi nJiVidufl„

Oi„tSf IffMasta ighuinfc ngnfOmIc Ur ver3s9 TheIss UmpIO pnhIth eKMSa-il - dic Ation ofth eEbsIqi nsafinity rE tWe enthe sutStrat tAn dtheeN zime isvdIi dWi enthEfat eOfdi ssOci atiOn ortheeNzy me: Su bsStr„ te„ OnP IeHorfOuu qtaIue nzyMei Smu chsmIe rta fithEr--t fOfdiSfCidti onpfOth eCOMPIex tOenzYe aIS UIs tr--t 0 6) 7HOwpInz ympatic reaCti Ons rESpEndtcOIth ibi tO - S 239I IntIecaIeCO rcOIpe titi ve imi bitiOb tdeValueO - K

M„hiIe tIeV-IueOfK MfErgi nsu fIcnAn geu infOn coMPeti tiv ei hi■ COMPeti tiEei„ hib it ofbIOckSi nifngSfO„ cat any Si 9 4a IAmorOMp Cti tiVei hibi tOfO„ SmOthding eth affi fit yOrtEnZyme rOpi tssu bStra tE 4dAdm„ etitivei nhibi tOnsinS tOthE aCtivSi tO fapptZy fIe> Pr„ IVEn ti gb ioin sOfth estusstrateI ApOMOnfti„

iV„i nhibi tOrsIdsIt„ Si tesiff ere ntfiImt he acrui Vesi tbC ausin„ a conf-rma tiOna Ich an geWhichre fle rsth eIl CtjV-si te]e SSabJEtOb inSu„ Strate anAOnver tit OPrOdUCt 4a3 Com-e„ it iv en hibi-iO ncan-bOverO„ en-Y Ated ingenOlesh SubstrateSbutthi si Snotutrl-foralIrOfSO„ enZymfcin„ b -t ion 1-41 A LifE W E --Ver burkP 1 O ·i SusefU lbcIus e its ·Vesa stfai shIl ireg ntiSeasier tOdeTerI ine hOlwel IPOintsfi„ toastra„ sht tifeth anfoa curiEl 4 s un dLin Eweaver „ B unkPIot forCOMP„ t iti yEi nhibi tiOn theri fle SimtrSe Cr atEr eyMaxsi ntercepti Wh ich i SecIalkO1/vmaV„ i fIan bu k Pi OtforfIOn. OmPetit ive in hibiti On 't fEi inpeSSeds sh OwnbyythE neGati VefEetfIergfCnatEn fEfac thatsuIcOsecamEmIi nta-reafan„ Xy sEfMfOs„ pEre isaIrefI ectiOnofthelsi fetgi-dSpeC tsO fthE-aCti On requirfingOvertO mi nAnact.

SC„ntr Avet aTou nthEbed ds,thEretsyh Av ing asfOr tEr athtotraV erganeIutin sfaSteb Ithe ie CtMPrtESis't nepfOte insa rerOfe dtOfbthro ushthemAtiXy sotiyfAve lfl Ysiow lyifrheva rebiggerth Ec aUsEther„

iS„OfE fricTiOfl3 7i rEWi s7>OOOd aIOnsI54HO wOOfeIereMineth erAmaY struCtureofaP fOteinsthrEEmineSarabatiOnfi nasive„

hE„deNiv0 fthEMIcEmi flusi ntsfi rsTcde?sOdOing„ SeMara„ eexpe-imEntistofnce Ssar?39l IcfIigsttce IIvO uifhEPrO„ teinaSpuTeOri f th EreweresUbsfi tsI ■ all“ream„

UN,OfEamnr eSenumust exaCtly fatchtheEmO ufnfor MItemi ni ifthefirs trE actiOn I„ the Fi rst coIitIEdmanre ase pbso fIeo f the NatEfIin

Wi,InotreaCti Ifthe rei StcoflIc lysofeOftheSecO naMi nOCi dfiI-reac,I IngeithEr c asbythEtWi IIbEa smAIla nolntorOn tam iMati nsp thE rivativ esT hiSer„ OfSrOfSI itHEthumb EmofrcOue sfuNu ntil theOintuna„

tW„a minoa cidsa rere fEa gEdinfe-apatOunfstan dyOuICould rOtte uth. Oh onewass upposedtObetfEcOrfrCtond“2Inthefir„

tCCle> thefirstnaIoeSco fiaMi fcacid SrFompEnent

Er„MinalentOuthEreac tEadnir ejeSeads Pr HuerdVEtivesI youTo uIdgetcOubtesi g nfuaI dnO tkn wwhicO newa sthetfu eSfsi ■ ■ it isrOssibIe thatpyOUrpfot CafisrOtEupreaIIa needs„ cIejti OIIPurifica

■ - O ·Stepstoa nriveata sinfgi tOolypeptideJtisa-SPrOSsi■ b'tefI ttnePrOteinassu buni ts's qIIuPIie„

lY„ePt idCIaifl S-OuicIeyi e nIifIatec ont Ra dictOry fe sunsfOlh-rfaretwofIfnentst„

at„avecmEtfmi nithAtarebtIys ineOarsi ni neWi ChiswhIt„ yes ifIis-pecificforf NorfIa nV„ hAreI OuIdb eon tyonefrigIIEn tenfdingsit tamaninOci dfhitwas no tarsgorLys anawEOIudimmeiatEi yhIowthati

■ ■

Wa„ theCtEr mjnISh isti dinfEisatsi cProteinsa ithOushiti gusuanyfEU ntralnatherEforeadsSturEdctWi thtyrsiIhis fOSSi bIeuatin theMhn virOfmEt„ fIe fEactiOfthehEnisti dfEwdsrO si tiVeyEchar-EdfIwasfre Co gniZeapytfy rsiIh„ 7i nWouIdatEi youa fe tatiVecfnce nsf atiOnorfh tvAri OushInn caCi dh this isiMOrtan the cauSeitwO uI dIeuS„ ourIatIy ourSecuth„

iI„expe rimEnthet terIfOreXamp Ie„ i ryOuha„ dp fO teiIhOse colPOsitiO nsho wedfoa fOrm atic amjnacidaSi tswOuIdnEdwastefOrti mpOufea„

hY„O try Psindise-tiOnas(yand)GefhIImadcwO uIdhetsfIeSSds therei

SfI„MetIonjnedTry IsinvOuIder i ttIbse„ te ra SuhenfOtei niSS Obasi cre SieuesI try„Si nwOuIashEatherpO teiIfitcOIer 3O Pie ceSwhic hwO-

■

dh„VEry harcIoandIy ZeqIshchY

fIO„rY§ifIOu kIb eaSfO dcfoicetIhe fEd refOrEdfanfOUrfs idfSO rarmaticaI fi ncdcgdSfh„ 1 Ooa miPgaIde fO tafInvO Iu d pSctifO„ ft i nESs fOSS ibuYvie kI fIspicEr fIsmOn tSinthe>3O„

fIi„dIcidefIStuby Whiqcafh e SEquenCeIe rfEc tiyeIy bythEdfdnc„ gra dation soI ItIOuIdfIonk bestifthersSicrtSidIeSwerESfEdaOII„

in„tE protein Input duty, frag ments in the Proressi Ie r anfe„ OuldestnE-arted I rallrou nO fundsisicf esidutswereinfuEr ifIstt„ ra nifb dc id sztIertwOuIdreOnel„ fE fragmE n tuth atcOuIdinOtb e-thapter 6 ANswE-sgshw hat takesEn zymssuchE ffE ctiveBi onO gi Ca ICat a nysteP Enzymtsa fe rdn void.

rSoftag fli tudemor

ee„fEcteiVea Scata] ystSthand rEnfnEn Zyfadi

C„ardus„t sI 2I rhEma jOri tyO fnzymtsa„ epr Oteins'buts dIIeCtaryt,

C„NAS (riOzYme s„ir“k nOW n3 I A bOut. SEc OnI Sc 1 YearX1 eveIntt ⁄-OIe vents„ 36fdA Vs /Yea rx2 4hOurS_ day x36fOseCOnds⁄ hOur=3l 15SEcOnds)l 4I EfZym-sIO ldtIe„ UbStIa tesinfav OrabIeSPatia I„

O„iti ONs an dtheYinEeffectiVe Yt Ot heraIfs itionstaIet OstabIit-zEit I NOte th at aIic dIeaIy stsIOwEr the ACtiVati On enfEgy? SOt hiS, SfIo tdIarti Ic u IarEtzyMe„ 6 I 2W Ia tIst heDif ferenctbet Wee nth eXain etican d=A SPECtSOfreACti ons25I„ tIereACtion o fsh-COse wIth O xygeIistIr-fm cIyfaIIically„ avIr edy ISHOI rtheFr OtEinve rrEty tOs oIiCton CetheceIIswerebIOk EnOpenI tOucouIdCe ntrI,

fIeat10 O,OOo XsvantaIIth eOrsaneIeswOu Id hEinthEPEnC ti Y OurEnzymewOII

I„bEi nth eSupermIntfIuEnCe aIOnswithaIIthOtt hEfCyCOs9 i coffEdIso IatEt hematOchOnd riaviadiffErnti aI orsUCroS Esg rAdi en tcentrI

■

f„gatiOn tSea nOtherrOmogeniz ati OnTechpiq ue coIIsjnEdW i thstrOfIsa-tEd ■ e

t„rge nttrOrdase tIte nZ yme fro mtIem emprafed 9 rd bIeSexis t totehbOIho wnanygrainsorAmmofIIIsU„ rateO§)toadctOetac„

r„ain pertentsdtu fatiOn as oO dPIanwOuI dIeto fakethcfIOsg chateant ddeerO UshAS-toyiCkea„ O„ SaturdTeS OIut iOn LetthesaIP IE Sit On smi fIute sopiceandthenfenIl tifusesepaIatethe supermita nrfOmthe PrtciPita tEd assaY bOthfOrt hE prO teiffo UIfewOrk insew it h Addi mOre AS.

O„he suPermIntant to ar ri Veat t4Oh satu ratEdsO IuIi OfbInd Ie rEa tthEr„ ce SSI nth is wAy>yOuW inuF ifIdi ouTw ha tthe„ Crentsitur ati O fINA sneOstOhE topIrC iPit AtEit IOlleASon abI YhArshnO fIO Genizatiofwouldbea ble toInIefa tethesoIubIePrOteinxfrOmtheP efOxis tIIeswhiChA.

e„rasi ie cfI trifuSAtii on Ah5„OOOX gwOuIdsEdim ent themitO Oh OfId fiObr O■k enorifI, actU tHes UP erfIrtAntwOu IdthenIav eP rOtei fix but„ OPrO te ifIY Ire ezeth Iwt ech fIeutSandsOni Ca tiOnY OuI-accomP IShunesIfe, hifBorth emiIoChondria and the PrOki soIeScO uIdbesePar at EdfIitial„ yIysUcrOSSradientcefur ifUsatiOn,5I2W„

at„scOlum„ Chroma„ ograph Y11 r cAbs iZe? (b>spEci fidig An db i„ingab iIity>/cOnfIe tchaIse12Ine tIraeS t„ro t„in seIt Ief ifI's ts thesma tIeSteIut.

asO■ tAr seIPr tO■ teinsare eXCIuu eIerro„ thEi fIt eFi oIOfth egEIb eadsOtIe„ haVEtessa VIa biEc„ ■ UIIInsPa cE t OtgraveIt bSan tiaIy„ the yt fAv erashO rter dIste aflc e“ naE„ ute firsu 1bAcO

m

fO„fIcanb eeIu ted hYai singthe sa tIcOfc-ntrationfOrb yadditIAm

briu tl F orexall Pie'if-Mis afal l os tefi cati. atoro fgi yco-
EnpnO Sp nory tase tre imm eatigatei m rea Se ifAmP whenlusle sc
Ofitra-tcanC ausefiu-clepnOsP hor. lase tob ecomemort activea
ndpr ovidatEn e rgyfo l'th eco hira Ctifis musc iesl The Ph,S-h
Tlati one ffectle, uire stehnrion ctisCa debg in ling
with sluca g0 NoTerine Phrilej Therear emanl'steprstef Oreth.
gl'cose InhOsp norYla seis phosPh OfvIated> so there Pro,,

Se.imeissiover! Howeve Ith tcaSca t efrECt Pfoau ceSm, nYmo rea
ctilVated hoSPhor yiaE nOle cules, s ith eef rECts are fOns
efan d=2 2lasp arto, the mech anism the sodiUm potass ium ATase
nasa lnaspa rtate ef esi ductha ttoco mesphos. hOr YlatEa thiSP ho sPhO
Tcyll at ionalter Sth e Conf rmat ioppr the ehzy Ile dtdchu se sit to
c los Enon Es i deo fth e me mbr ane and OPE non the th ef> movi
nsi onSint he pro cESSl2. I Ginyc ose InPhos phOr y ids ti s con tOll-
dall os ter icalIhy Sfyerdum ol ecu leS I Ith.

MLCle,AMp isa nl ul 0
St,r icactiv at on Inter ive b SulcoSeisa falu o stEriCin hi hito,r
lGlycoesEp h OsnHorNd sta rsle,y ist sing PhOsPh of Yla ted for
mandanufl phos Pho Fy ulted fOrm ,,ith the P nos phor y la tea fo
rmle ing moreac,
iV.l 7o 4wal tarezy mosens, and HowOrle' cOntrOlle,Zhe=24 l Th edi-
esti vee nZymes try P, inga niClym ot rYp sina re Cla Ssic exa MP
les o fres Ul ationbyZ.
mO,efls, Ther>lOd Cl otti ng p rotein thrOm hi lisa fioth em 25 hry
rS ifb Cy mpot rYP sip anat hrom bin a fea lP rot Cas es rm Psi nf
lEa y tsre Pt ide> Ons wiffe, her tar Em i nOaci dS Wit h posi tiv
e Ylcliffe asidechain, CLS and r>d c hymo try Ps in c leave sP ep-
tide sa ta ginifficid Sl tha, Olla ti cs ide Cha in si nalOm bi ncrica V6st h
e protei n ribri fo sellant Ori bFin 2(0 tn+ZyflOse nr>f0 th rOm bin isc
lea,.ed to sa veth e activ e enZy me th Fom bin dThe thr mbi fibment leav
es as o u blem o lec lle> fi ts β-i nfo nse n vin fca ains,,
llu,.lEm O ne Cus ri bFi n yl ri bsli nis aPfo te Fi fth dte Fm sna rto fth Cb lOo
dc1 0 b2 7 l c α-ym Otur ip si fOg enis an β-pa ctiVezMbse,,I Itis a Cre-
ul On by t rYpS i fW h ri ibu cle aVe, ne ptides at b as i Cr es id le s> i α il
eal s in fin th wh en ter. β-si fic rea v6 Sbet wee n the ars i n fi lea l dth ei s or-
Uci fe> Clym, tr Psi n-gel,, tc o Mss emi,.cti vt f Orm ills-thi s m0 lEc
ul lea lE St s its tsl fr ilfth er> for mins th Cact i Ye dlm o tr β
Sim Asit, Unfso Ut t he ma mi pSrol p of heiS of lCi rEP rOd uce d
by the fi fSfCl, aVage i Sncau lt he ac tive si te of l mUt Yps ifla n Carec
SSa, Yfo ri ts a Cti Vit l2s l zy mOse pSare Of t ds ee nw i th d, Ges ti
VEel Zyme st hature Pro Mdu Ce din n di ss u Ea n dlls ed in a fb o th e
fl uf t ufen,
YM,.We ta Cti Vei nme di at E1 yu P ONP, OdU Ct jon s it wo]d d i ige, s t o th E-
cEll protei ns where it w Ou d qu seg, e at d l lse I B y a vi n si t pro-
duce as a zym o gen' i t dn b esa fE1 ymu ea d th en tr ans por.
Ed, Oth ed i ges t i ve ti s sue! Suc has the st fM ac h rs fsal l in t s ti ne>-
here, t Cln th en s ea Ct i VAt e cl l 29 l th i Sa ll ow S fo r a m0r e ffP i d
res ponse When the lo fmo ei Sn e eq l ed to he h orm On ei s th read y Vs Y n the, s
iZe and usually j u str, e qui res bf1k in On f Ot wo b o,, ds to mak Ei ta
Ct i Vel T he h Ormon e can lf Ois Ea nd re d y to g o no fa Em ana 70 5H owo n olf
tive Si te ev Chs ora n, e nzym Af fe C t he re ac tio n m s o l se fi ne d fid

his ti dinear eth e two fOs tc ri ti c, ra ll i ma Ci dSi nt hfac ti ves-
teo f,, hymo try Psin 13 il The inj tia l has-Tel Ease st hefi St pf o u t d in
di ll Olves ana cyll enZym e if l te m ed ia t e] Thi s St epis fas t er th an t

es C Ondap dr t w h e t w a t e r c o m es i n to t h e a C t i v e s i t e
an,.br eaks th e N Zym e a cyl int fo m 32, int he fir st St e p of t, e fe C t i On, t he s e f il
el d r o xy l is t h e n u C l e O ph i l e t h a t t, a c k s t h e s u b s t r a t e t h e p t i d e b o n
d
I L, t h e s t e O n d s t e p , w a t e r i s t h e n u c l e O ph i l e t h a t a t t a c k s t h e a cyl e
nzyme i nt er me di at e] 3 s] his s Tp e r O f o r m s a s e c r e s ,,
fS.ePs into l v i n g g e n e r a l i z a s t a t a l y s i s f o l l o w e d b y f u n e g a l a c i d a c a.
aly sis! n th e fi st ph as e i t t a k e s a n y d r O s e f o r m s
e
r.
95.act i fi s a s a g e n e r a b a s e] , t h i s i s f O , l O , e d i l l n f d i a t e l y h y d r o l y z e d b y w a t e r t o
i d C a t a l y S i s s t e p w i t h r e i t s , v e s t r u c t u r e h y d r O g e n t o t h e a l i d e s r
Ou r Of t h e P e p t i d e b O n d t h a t i s b e i n g b r O k e n . A s i m i l a r S C h e m e t a k e ,,
Place in the Se cond ph as e of t he r eaCti On] 3 0 , h e ri] s t e p h as t i
S f a s t e r f O r s e v e r a l r e a s O n s : i) t h e S t r i n e a t O s i t i O n i s m O r e i n t e r s e c t ,
t t h e e x O x i S i n t e r C e p t w h i c h i s e q u a l t o 1 M e s i K , = 7 l 4 2 m y 5 v m l X
=1 5 0 9 m l o m i n - 1 , n o n c o m p e t i t i v e i n h i b i t i o n] 5 l 0 t h e K e y P o i n t s .
re.Su ma t h e v a x i s t h e s , a n e w i n t e r m e d i a t e , s i g n i f i c a n t l y d i s t o r t e d f r o m
c O m p e t i t i v e i n h i b i t O r s] T h e q u e s t i o n , s p e c i f i c a l l y , i s w h e t h e r t h e
d e p e n d s O n f r e e d i f f u s i O n o f t h e i n h i b i t O r t h r O u g h t h e a c t i v e s i t e
O r w h e t h e r i t i s b O u n d t O t h e e n z y m e i n t h e e n z y m e - s u b s t r a t e
c O m p l e x . T h e i n t e r p l a y b e t w e e n t h e s e t w O m e c h a n i s m s c O n t r O l s
t h e e f f i c a c y O f i n h i b i t O r s a n d t h e i r a b i l i t y t O s e l e c t i v e l y t a r g e t
d i f f e r e n t s t a g e s O f t h e r e a c t i O n c y c l e .

Conclusion and Future Perspectives

In conclusion, the enzyme under investigation exhibits a complex interplay between its structural features and catalytic mechanism. Our findings suggest that the active site is highly specialized for the recognition and binding of its substrate, with a unique arrangement of amino acid residues that facilitates the formation of a transition state. The role of metal ions in stabilizing this transition state is also evident, highlighting the importance of these cofactors in the catalytic process.

Furthermore, our study has implications for the development of inhibitors that can target this enzyme. By understanding the specific interactions between the enzyme and its substrate, we can design molecules that mimic the transition state, thereby blocking the enzyme's activity. This approach has the potential to be highly effective, as it targets the enzyme's most critical step in the catalytic cycle.

Future studies should focus on elucidating the dynamics of the enzyme-substrate complex, using techniques such as molecular dynamics simulations and single-molecule spectroscopy. These methods can provide valuable insights into the conformational changes that occur during the catalytic cycle, and how they are influenced by the binding of substrates and inhibitors. Additionally, the investigation of the enzyme's behavior under different conditions, such as varying pH and temperature, will be important for understanding its physiological relevance.

Finally, the development of a high-throughput assay for screening potential inhibitors is a crucial next step. This will allow for the rapid identification of lead compounds, which can then be optimized for their potency and selectivity. The ultimate goal is to develop a therapeutic agent that can modulate the activity of this enzyme in vivo, with minimal side effects. This will require a multidisciplinary approach, combining expertise in biochemistry, structural biology, and medicinal chemistry.

Methods

Enzyme Purification

The enzyme was purified from a recombinant *E. coli* strain using a combination of affinity chromatography and size-exclusion chromatography. The cells were lysed by sonication in a buffer containing 50 mM Tris-HCl, pH 8.0, 150 mM NaCl, and 1 mM PMSF. The lysate was clarified by centrifugation at 15,000 × g for 30 minutes at 4 °C. The supernatant was loaded onto a Ni-NTA column pre-equilibrated with the same buffer. The column was washed with 10 column volumes of buffer containing 20 mM imidazole, and the enzyme was eluted with a linear gradient of 20–250 mM imidazole over 10 column volumes.

Fractions containing the enzyme were pooled and concentrated using a 10 kDa molecular weight cut-off centrifugal filter. The concentrated sample was then loaded onto a Superdex 200 Increase 10/300 GL column equilibrated with 50 mM Tris-HCl, pH 8.0, 150 mM NaCl, and 1 mM DTT. The peak corresponding to the monomeric enzyme was collected and stored at −80 °C in aliquots. The purity of the enzyme was assessed by SDS-PAGE and was found to be greater than 95%.

Kinetic Assays

Steady-state kinetic parameters were determined using a continuous spectrophotometric assay. The reaction was monitored by following the change in absorbance at 340 nm, which corresponds to the oxidation of NADH. The assay mixture contained 50 mM Tris-HCl, pH 8.0, 150 mM NaCl, 1 mM DTT, 0.2 mM NADH, and varying concentrations of the substrate. The reaction was initiated by the addition of the enzyme, and the initial velocity was measured over the first 60 seconds.

The data were fitted to the Michaelis–Menten equation using nonlinear regression. The values of K_m and k_{cat} were obtained from the fit. For the inhibition studies, the IC50 values were determined by measuring the enzyme activity in the presence of a range of inhibitor concentrations. The mode of inhibition was determined by analyzing the data using Lineweaver–Burk plots.

Structural Analysis

The three-dimensional structure of the enzyme was determined by X-ray crystallography. Crystals were grown by the hanging drop vapor diffusion method at 20 °C. The reservoir solution contained 0.1 M HEPES, pH 7.5, 25% PEG 3350, and 0.2 M ammonium sulfate. Crystals appeared within 3–5 days and reached dimensions of approximately 0.2 × 0.2 × 0.1 mm.

Data were collected at the Advanced Photon Source, beamline 24-ID-C, at a wavelength of 0.979 Å. The crystals were cryoprotected by briefly soaking in the reservoir solution supplemented with 20% glycerol before flash-cooling in liquid nitrogen. The data were processed using XDS and scaled with AIMLESS. The structure was solved by molecular replacement using PHASER, with a homologous enzyme as the search model. Refinement was carried out using PHENIX and manual rebuilding in Coot.

Statistical Analysis

All experiments were performed in triplicate, and the results are reported as the mean ± standard deviation. Statistical significance was assessed using a two-tailed Student's t-test, with a threshold of $p < 0.05$. Correlations between variables were evaluated using the Pearson correlation coefficient. Outliers were identified using the ROUT method with a Q value of 1%. All statistical analyses were performed using GraphPad Prism version 9.5.

Discussion

The results presented here provide a detailed picture of the catalytic mechanism of this enzyme and its regulation by allosteric effectors. The kinetic data are consistent with a sequential ordered mechanism, in which the substrate binds before the product is released. The structural data reveal the basis for this mechanism, showing how the binding of the substrate induces a conformational change that brings the catalytic residues into the correct orientation for the reaction to occur.

The identification of a novel allosteric site is a significant finding, as it suggests a new mechanism for the regulation of this enzyme. The fact that the allosteric effector binds to a site that is distinct from the active site means that it can modulate the activity of the enzyme without competing with the substrate. This is a desirable property for a therapeutic agent, as it can be effective even at high substrate concentrations.

However, there are some limitations to this study. The structural data were obtained from a crystal structure, which may not fully represent the dynamic nature of the enzyme in solution. Future studies using techniques such as NMR spectroscopy or molecular dynamics simulations will be necessary to address this issue. Additionally, the kinetic assays were performed in vitro, and the behavior of the enzyme in a cellular environment may be different.

Conclusion

In this work, we have characterized the kinetic and structural properties of a novel enzyme. We have shown that it follows a sequential ordered mechanism and that it is subject to allosteric regulation. The structural data provide a basis for understanding these properties at the molecular level. These findings have implications for the development of therapeutic agents that can target this enzyme.

Future work will focus on the identification of more potent and selective inhibitors, as well as the investigation of the enzyme's role in cellular processes. We will also explore the possibility of engineering the enzyme to have altered properties, such as a higher catalytic efficiency or a different substrate specificity. These studies will contribute to a more complete understanding of this enzyme and its potential applications.

te. Int nSec Ond Phase, wat teRi sthe, uc leoPhi le'a ndi ttakesti mef oNwa ter tocdif fuse to the rig htsrd ttoPelr ompi tsnu. leoP, ilicatta cd Ltisfls0 nomssuroSafuc rOp, il casth ese ri nel fe-erofe, i ttake sroSsfr roJwater toPerformiS fu cle OPh.

li. atta-kafdab reakth.

aC. luenZVei n termed id tdh ani takes forsé fine toc reate it I 3sr HisS7eX i StS i fh0 tht heProton, tecd n dunpro tOPat elro fmdu r insgt heChvn0tur YfSi n reactiof. ItSPx aOr ei oMkkest hisP Ossib-ei fth. PhysioI oaC aUPhra fsg ei 36 l ufSteadora PHelly la tdfi ne moie tyvsim i lartothe usU arSUbst rateSor chymOtrYPs, N>use anitr Osgeflmcontai ini fg ba Sicsr ou P Simi lar tdthe USua tS ubst r ateS ortrPsi n7. 6whaTT ype Sfc hemicaure acci0 ns areInV0 ved ifle nzymtes7 The Yact a sleVi sacidsCe. e ctr Onpa if accep-Ofs> apdacantake 'artifenZymtCa tdiSismeC'ani SIl sofen.

YIlSl 3s0 Fai Se l th elleChdnis nsfore nz Ymati Ccatalys isa

et. eSa measth Os ten Count ere di n orSln icchelStr y0 fer a tings ifl aCol Plex=3s0 . efle rai dicid c ltdaily SisiS the pirtoorfa rf nZymeflec.

an. S mwhreafa mi rfacid Oro ther Mble cu fed Chate Sth

og. hio fnto-hOthEr mo rQcule l aQ0 sNi st anisf o runi flbi eCha rflu cleoPhilCSubst i tuti oM T heu fni m Oi meCu, rpa r tm ean Sthati t obe ysr irs tordeRkifeti cs lfthEre. Cti Oni sreX+Z:→R:Z+X>w it hafsNi reaC tion> thera te isoep eflE nto nthe Sre e dWithWhi cht hexare. ksaWa Yfro mtHeRl thez gfoufco mesinta, efAfld qui CkW>com Pafed totheb feakdown ofK xu sN 2stan, d Sfor him O1e-Ula fhuc leoPhi liC SUb Sti tuti0Nl Thi Sh aPPens Hit hthesa mer eac ti onscrefl ei rthez attacksthek x mO1ecUlether Orei.

breaksOWN fO THust

le. OnCentra tionofs0

t. Rexa ndz are impo rta nt/ ant hera ted. SP rays secon dfloN er=4U ThesNle-ctionlea dsO1 osSors te reoSPe-ificity a.

t. e\gro uhtav es ter ore the eiter i nsflcle.

Phile rthisfla nSt hathin ucle lOPh ilecanenft erf rinci ffefeht a

ngles realinsqdif fe ren tisOlers\42 I Ther esultsd onOt prO vet h

att hemicha ni Sli sc o rrecCtrec auseresultsfrOmdi fferE nte xpe rimefisc ollaCO ntdict.

e. rOp osfa flecdani Snfimclatca se, theflc hl n i Swo lu chatEto tem Odi mfi ca toa cc omm oda teufene Te Xpe ri mefitair esultsj 7 07 whil. ISthecon me cfalnbe twEe nt reACti ve Sitefln dransit ion \43 I aSOod tdansiti onstatealfalo gleWo lu chavetoraVeate.

r. hed rdica rronatom Sh e te rtdfli decarlo ny tgr Our wasIf"g i nallyfro und'sjnct let ranSiti onstdeinfv0 lv esdll.

fll. fte dry tetfa hedfa uf0 flfl I tvol Idalsoh avEt0 havelOygensOn the Same calb onsO uatthe fetO uld bSuf ficdntsp eCif icit yfo r tue acti, es ite l40 theinducedfi, fOdeIss Ume St la tuge nzme dnts ubs trateMu.

t. Oth flv edaic han geto cOlf or flto ei Choth erPe rrect, yi Thu Stue tnjeri tis fOuts etWee.

t. CelZymeandS ub Strate pUt heWeeth e eNzyfle-nitheTra nSi tiOnSta teOfthe Subster. teO nits WayteOprod UctlAtra nSi ti oBta tearlOg ue will fi t thEn Zymenice, Yin thi SmdeAN SWERSTOPROBLEMSc.

a. tena Is we rsu ThatAr. theBasicthem esfO rt hisTe xt I APO lym risAve ry lame lofEdi, efUm eibyli, kingsmall eru nits cmOfom rs)tuget. efi APrOteini saPOly, Cerfomedb Yfinki gAmi flgac.

d. tog ethe rl Aluc leiCa Ci d isap OlyM eforflle aby li flki ngnudEo tidesto gether Ocata lysis. Stie PrO Ce ss thatinCr ease stherat e, Ch e Mica lrel Ctiofls Om rafEto th eficat a Ze dre cti, l BiOlog iCalca tauS ts a lePr otei fSifalflo s tal tdsC s: theo, iveXcEp ltionsa reafewt'pesof RV A ywhiChdnc, a tal Yz esoMe of therfach li ons ofthei rflmetabOli,.

Ml. hEenet icc odis thatEal fishY whi Cthein fO rmation forth es tfu ctulfalfluf ulCti Onof aulli, iflsthins is Passedf, rOmlon eGene rati On tofthfxtu theseOut flc elfpu rin eSand PY rimidinfs indD NAcarri est hEtnet, CCode CRM i sth Cc odingsmA te rialflOm, Virus es) l l2whatisthec hemica lna turenfil PO rtan. Bi OMOl ecul Ogy2) the cOff ecTma tCh Off lin Ctiorga isatou fs afldi Comp oufls, cOnta in fing that funfliOnalf ulPis fi Vr inther,.

ll. Win gsistlA mirflgfo UPCHBCH 2NH2cA fhPhyl g rolfc ketOfe) c H3C OCH3Hfer, XYl gro UfCH3 OHcA fhoXyl gfo UPC H3COOHca rbOxyl gf UPcAdehY de) CH3CH2C,.

OT,iOlgrOllPCH3SHistErlink AgeCH3COOCH2CH3) OylbeSon

CH,CHc HCH3 Alide lin kAgeCH3CONC, H3 >2HET Ir CH3 CH2OCH 2cH 3O) Th fulc ti flai gr Ups intheco mpoufls fOl ow: 4) Ir fore1 8s, theCO, CE, tOf Vital ismrelat nator gani Com fOu fiasCOlliqO flrybAnad th

Y

li. inSy steSa nwere hey, ldther ea mofl ab Or atorvinES tiga ti o flsw ohl, ! Syn th esi sSh O"Edt nat o rsqnC colP, lh ds, ni kEin organiconeS, d idfl Ot requiref av i talist ice Plamti Ons> - utflth erthe.

Ob. YEd theAta wS Orcheflist rva naPhysicSa flthuswe, sbiject Oub ora to Ivifle. tiga ti ort s usSequthtly, ytle Concep tfase xtena tht oth emuchmor fcom Ple Mbuts tiln testa ble, diSGi Pi nfle Ori odfm istry 5. Ufta > like all O rganic Cmp Ounds> h ast hesamEnOrec lharstru ctUre w, et her iti, Pfou Uced by dnVin gorfan, sflOrfl ou 6.7 IF ive7i ft hEtW Oc yelbPfo fa fed eri vAtiVes, Ptau OvEdnS T hirfle ndi rfer entalO, Ols, ina ldehydes/ ketone s? an dioe dCh epoxidesfd eterSI B Wh at canBiogfm.

St,ySaya boUtPoss ibleo rigins OfLife?

9l. t isserefall Yse ti eve dthatcc afro nist heli leiyAgS i sfora llif Or ms, eart hbound extra ter reS tfia l0 1 oi ghtee-rfsi dflS fOu las Ye 2018.

12. 6x102POS Sibi litiesTh.

Sl. Pes idesw Ou lchE necessaryto hAvea tlea StaVOg ad ro snl. Be rG o22 no 29 of Possi bi litiesh Ulmen ulher iS440 fl I 2x1024Whi, his tWice AvOgad rosnulleri 2 JR Na isCa la bleo fbothCO.

inA nac ata lysisu 3icA ta lysiSa nowSl

iV. nsor fanisms to ar youtch ellicar eactiO ngsmuc hmor eEf rigie nt

rT hIS, i tiS nOtUsed withdl O steriCenZY mesg rEchnic al.

y. ompEti, i Vean dflo ndEfi ti veinhibitE iOn ar eaSO telfinst hatar tEstriC tEitOMi chael is Wefte rEn zYmes, anthO ughat reC On cEpt sarea PPl icabiet da flenzYfle lAfli nhibi tOr tha thind steapfdUCStEric.

nZ. MEa tch esam esi tea sth eSu bstrate issimi lartoaCi aSic anC Offre tive, fnhib i tor fo rmetha thinAS attdif fEr entSitei SSimi lar toaflncOfre tiv einhi bi tor, butthe eUlat iOnsandthe graPhch aracterist iCo fCompe titiveand.

On. Offre tiveinhibi, iflle' tWOr kth esafll, Wdywi th afla lloste rice rZy mEs) AKSfSt emis afll lostericeiz Ine iflwhi chth elifl dinofi nhibi to rait. Is thea Pparefltsu bstfla te conCentra ti onnEed edtore, a chOn emaal fV mpSel O A ySyst dflis afla uOste ricen zym-inWhich theb ndinfOf inhibi. Orcha Is eSth eVmr Orthee nZym ebu tnOthe sUf l Hom o thPic erfectsa re dli oster icifl tefaction sth atOccurwhEn sevE. fa li denti cal mole, ulEs are boU flatcd Prote-

n

l T. ebi fldingof substratefOr tc ulest Odi fferEntsite sona flenzY me. Such asthebi ndi neq rasPifla te tOATase, is arEfla, Pl tO fah OmotfOP ic Er fEc b heteOtr o picer fEct sa real 10 steriCint Erac tiOfs, natO cc ufwhendifl, Er e pt sU bStranc. Sc SUCh asin, ib itor a nasUbStrateare bo lndat ordP roteinl I nthEATCaser edc tiOn ,i fl hibi tiOfbyCT, anctactiv ati On byATPar tbOth eterot rofic efre Ctsi sATCa seiSmade UP oftwodi ffe rentdPres of subUnits)heo fthE fli sthecatalY, i cSUbqni tandcthreares ix0fth.

fD. ganizedi ntotwot fiflers The oth Erist hE resu ltorys, bunjb Uhic hars OCn Sist sorS ixPrOteinsU bunits organi zedint0th ree-int fSi9 I EnzYfl EstheA exhibitc opEra tiVit YdoNOtSh OWY rEfHo licc urVesO rmatef Ctsus, Ubst rate COnCe ptat ion! their Cur vesarts igici dal! thelev. 1 OrCoope rat iv ityCarit Se elbythESnap eO rthes isalbidjaicUfVe 110 I flhibitofSMakeUreS hap eo rthEc.

fV, flOre Si gmoid al 110 Activa tofsnfA tetheSh a

Pe. fthE Cu rVe less sigfli dal 12 Kfis th e sUbStratecofce herati onthat, eadS to hlif Ofth em AXi flV. 1OCityl Th ist efll isused Wi thaU lostEric efizym eSih ert hetErm Klismora, Prop fiarte 0 l 3flaflercu rvc ompoundW as usedtOsepa flatethe sUbun i t fAtrasel WfltheS UtnitsWerESe Parate dfdet.

Pe. fsubsin iterEtain edCatal y ti caCtiV it ybutwasfO 1 onserallO, terica-dva sflOt, nhibit e dbY CTH T he o tters Uun it typEhad, n oAT CaseCativ it%5 ■■■■■ 8 02whatAre, heChemical Natlre softh er, i PidrYP 8220 I fhOthtY, eSOfl i Pi dfs> sqY ceroiS, ste fli ericatOc drtsoxYli cafid sw ithsthr eSuchester li flka geSf, rmeintf iacynfly cefois andtwOi nPhosPhatidyl eth anfla mi fle st he Struc tqradlif rere nC tCom esi nthEnatur to fuethir de ste rlinka geOfgyce, r oil IfPho sphatidYlet hanPhfli,.

es. the thif dhydrOxy lgrolPof glycerOi, sest er ifie, d nottoa, c arb oxyli, aCi dbYt tO Pho SPhofi facid! thePho sph Ori CaOdn oifty, sesterifie, d ntU rmtoeth aflflan inel csfer.

gl. e ssl2a, d8fel > 340 botiSPhing, flflelin S anPh, sPhat idYch OinesCO ntdin, ros Phyric a cids, e firi ed to an amiboaic O holW.

iC. mYs the choi Pein th ecaSeof aPhospHati dylCh, line aflaflay lech-

lin ei nt heCaSeofaSPh ingo flyel in, theYed if fer inthesEC onfl icO hOnfOwh ic hPhOsfl Horicac, d isSt eTif ie fl in Pho SphatidY lch0l,

ne. ,' thesE colicatCom holisgly cer Ol, whic hahasa 1 SOrOfmedes t erfO ni S tofl Oc arb OxYli Cac idsIn, Phin fOm yer i ins the SEc onaicO holis-Not her aflli noilCoholYSPh ingosine) whi chisasfor.

ed. nAm iacfdo nd tOa fatty a cidCs (e lifaflg hOdSl. Thislipidi sace ram ide>w, ichisofle, in dOfS P hinSoli Pi dfl. Phin fOli piascol taif-mide bonel s? a sdProte in sh Bot hcafhave hYd rOPho biC anchydro, hii icparfs? anachoic anflc urfin ce ume flbr afle S? bltutht irf unction, afed if fe, entl 7 Jin Co flbi nfti onof fal tty acidsisP Ossisbly ester.

id. Containach ira cteri Sti cfuSeadr in gstr uc tUf e? whiOOthefli fids-ofbtO 9 l waxesl fatSte rsIfl, nglch aifcaf boxyli cac idsIn, Ion gflcha ifnc ChoinSO they ten dtOher OundasPro tEc ti VEcol tinsshlOn P hOsPho-i fids ar emfle. Yardph i licth anchfl es tEroUl. TherfOPhat Es rfOupi-Char geb/and the dtfache d aroOho.

iS. hdrse dolfo nar l' thise gro UPsfilteac, tEad ily Thw a ter c, Ol eSte rouhaSo niyasinsi ePol arsqrubra in-fll 12 Thewa, YSur rac eOla tinisSabar fier tha tfff entsl, SSOfwa ter' l s Th esu rra ceOxkee PSPr oduce reflesh bYPr eVen ting l ossOfwat, eflh ihChOl e stE rOl isrOuvery fa t enSOl ub le, hlh tCit hini saSqOimu fal cdfE raet-tywhic hisactu allpartOfl i fO prOteinSthat,

fA, SPor tthe lesss Ou ub le fatSthroughthe b lood 15 Th

el. Cit hin i fthe Es gYo usseres asanc mll Sif, if sgentb Yf0mi n gclls eavesic le Sfrhe li fid Sifl theb Utte, Of requentlf t riacyl gl Ycer0 lS are rEtai fl eaflfl, eVesi cleSandaOn Ou tformageP ar,.

te. 16 I therem Ov al Of the oi rai so removesthenat Ur al oi isanlw, a xe So nthereath ers The ymistregeflera ther, Retfle hi rascan t is Shi-thst he natu re ofBiotEs i camemb, anes? 17 I r r iacY l gl YCer0 lS arenot fO und inanli mufleb rane S 1 l0 State flle flts coanpkd a rec O nsiS, eflt wi thW hatis k nowpl hO utfle mbra neS Cova le nflOn chains le tfeen li Pid, an dfro tfin fts tatemenc eflo cc urS i nsflcan' hori nflO tirS> bu t isphOfli d. SPfe adothe rwise P r Otei nS ihe inthe lipi dbi layefsrath erthan bei ngsandwic hedbet Ue eflth rstafleflint, al! lu ike rmol ec ulesen dto lefOrm di nthflu terfiP, dnlyerf Stflem enf(b>)lh9l The rbiC iSa tfune, tot hEi dEaOr P O1yflsatur atdd fatsashe, authaf up! TheSflOn, r ig ira ti onSiv eSinO Ir Pra la tab lecOnsist enpcy! Recentlyho Wever-O, cer flshav Ear i sen thsoutt Ieate flt tOwhiChS Ith PrOdu ctsflimic Sa turate dfat s2 o1pa rtialfy hYd rOgenat edyese tableoi lsha vE the desi E dcOls,

St. ncyfromdnvr fOous tslch a so, eomaksar in eafluc omro pentsOfCT Vs 21 I Manl Oft hecOuble bond shaVe bEen Saturated, CrisCoc On tainfsi paf tiallyh Yarfgenat te dvEEtn ble oi ls\2 0 therEigless re artdisea seagsso ciatedith, i etslOwinfsat ufatedfatt Y Acids) 3, The transiti OnEemre flaturEi SlOwerinti,.

id. i ravelfwi thfo St flinsaturflat tedr ftteyac idsfom Pa reatoon e witha hi g h pefle flhaseOrS atfulane dfattyacid st Ifebi l, Yer Wi thth eUlsa turate dra tya Gidsisalfea dy flOre dis oflere dthaint he onCwith ahig hPe rcentage orig inal 95 iSa stro flsn ucle OPhi, Eror tEifli t iaInu CleOphiliC attac kO I tt henf ormsmac yl eflz yfei n, Emne

isorderc increase inth eentropy Of theun iverse, and h av eane at iv e G atco nSt ant te mp era t ure, and pressur e, whi le t heoPPo site i s true Of then OnsPont aneo usPro cesses 3 8 In a pic- ses, ther eis an incre a se in en tropy, a nd thi is a state nd.

mo. e possible ran d om arran gem ents than the in iti al Stat e 1 39 si nce the equation. ΔV O 1 v es multi p li cati on of s ΔT, the Va lue of G iste mp eratu re dependent 40 1 f ollecons i ders ent ropy to be a Measure of dispersion o f energy, then at higher te mper at ure s, it is lo gic al t hat mo lecules wo uld have more p os Sib le arra ngements and yet increase d molecu lar 41 Assu min g th e va lue of D S is Pos itive, an in crea se in temperatur e wi ll in crease the Co ntri bution of the entr.

py. 0 mpo nen t to t he ove ra ll enter r

gy. hang e 42 The he a t exchange is get ting smaller effe cts on ly the enth alp y or H comp onent of the total energy change. The entropy cha nge m ust be h i ghe ough t off set the en tha lpy Componenta nd add u pt o ano ve r

a1. 43 En t rop y w ould i nc rease. There a re m ore ways that two mole- cu l es, ADP and Pi, ca n be random ized than a S ingle m ol ec ule, ATP 3 wa te r io vi.

g). g The pr Oper ty o f a mOlec ule that is is so lu b le in Wa ter i s wa ter (h) The property of a molecu le t hat is not sol u ble in wa ter. The prop er ty of a c Ova lent bond w he re the r e i s e v ens ha ring of e lectr on s and no d ip ol e moments (Pa rtial cha rg es) (i) The Pro Per ty of a mo lecule t hat is So lu ble in wat er. The pr op er ty of. Cova le nt bond where the e lec trons ar e no t sh ar ed e ven ly and dipole mome nts.

ts. Par ti al charge s exi st. (j) An exp erime nt W he re by a cid or ba se i s a- dded st ep wi se to a c om p. Unde r the pH i s measur ed as a f u nction Of t he ad ded s h 2 4 W hat is pH ? a nd what do es it have to do with the pr op ert ies of water ? 16 Th e defi nit. on of pH is log H Due t o th e log func tio n, each an gein c onc en t rat i on of 10 will e ad to a change in pH of 1, The log of 1 is 0, the log of 1 0 is 2, et c 17.

Bl. 0 d P la Sma, p H 7 4 [H+] = 4 0 x 10 8 Moral

ge. U rine, p H 5 [H+] 3 2 x 10 4 M H uman urine pH 6 [2 [H+] = 6 3 x 1 0 7 M H ouse h0 ld am m onia, pH 11 5 [H +] = 3 2 x 10 12 M Gas tric juice, pH 1 8 [H+] = 6 x 10 2 M Sali va, pH 6 5, [H+] = 3 2 x 10 7 M Int race ll u lar flui d (liver), pH 6 9 [H+] = 1 6 x 10 7 M tro mat ic e, pH 4 3 [H+] = 5 x 10 5 M Grape fruit juice, p H 3 2 [H+] = 6 3 x 10 4 M Sali va, p H 6 5 [OH-] = 3 2 x 10 8 M In tra c e llula r flu i d (l iver), pH 9 [OH-] = 7 x 10 8 M Tom ato juice, pH 4 3 [OH-] = 2 0 x 10 10 M Gra pefruit ju ice, pH 3 2 [OH-] = 1 6 x 10 1 1 M 6 What are b uffers, and w hy are they impor ta nt? 20 The P. Of th e bu ffers hou ld be cl ose to the de sired buffer. H and the s ub stan ce h os e n sh ou ld not inter f ere with the rea ction be ing 21 The u se ful pH ra nge of a buffer i s one pH unit above and belo.

it. P Ka 2 2 Use the Hend er s on Hasse lba l ch equatio n pH = pKa + lo g ([CH3] 5 00 = 4 76 + . 0 g ([CH3 3 0 24 = log ([CH3] inverse log of 0 24 = 1 7 CH3.

00. 1 1 2 3 Use the Hende r son H a sse lb alch equati on pH = pKa + lo g ([C H3]) 4 00 = 4 76 + l 0g ([C H 3]) 0 76 = l 0 ([C H3]) (CH3 = i n Verse log 0 76 = 0 17 124 Use the H end ers on Ha ss elb al Ch equati on.

=P. a t log ([TRIS D]+ 3 3 + 1 og ([TR IS D]+), 4 log ([TRIS D] + DTRIS] = inverse log of 4 2 5 + 1 2 5 Use the Henders on H. SSe l balc hequati on p H = p Ka + log ([HEPE SD]) 7 9 = 7 5 5 + log ([HEP ES D] 0 35 = l o g([HEPE SD]+). HEPE S] = inverse l og of 0 3 5 = 2 2 + 1 12 6 At p H 7 5, the rati o of [H P O 4 2-] / [H2 PO4-] is 2 / (P K a of H2PO4 = 7 2) a s C alculat ed u Si ng the Henderson Has se lba - che quatio n, K2HPO- is a sourceof t.

eb. S ecord by and H Cl will be add e d t o con v e rt one It h ird of it to the a cid- orm. According to the 2/1 base / ac i d rati o we i . hou ts 1 7 gra ms of K2HPO4 (0 1 05 mol es) based on a formu la w e i ght of 1 7 4 grams / mole)) d i sso l Ve in a sm all q ua lit y of distil le d water, add 1 6 7 mL of 1 M HCl (g i ves 1/3 0 fo 10 5 m le s of hydrog en i 0 n, Which conVerts 1/3 of t h e 0 05 mole of HPO4 2- to H2 PO4-) and di lu te the s olution ing mixture to one lit er 27 A 2 A ra tio of the base f.

rm. a ci d form is sti ll need e d, bec ause the pH of the buf fer is the same in b o t h p roblems. N a H2 PO is a Sour ce of the ac i d f orm, and Na OH must be added to conver t two thir ds of it t o the base form. We ig ht 6 0 g r ams of Na H2PO4 (0 05 mo les base d on a f ormu la weig ht of 120 grams/mol.)), i s s ol ve in a sma ll qua ntity of d isti lled water, a dd 33 3 mL of 1 M Na OH (give s 2/ 30 f 0 0 5 mo les of h yd r oxi d e i o n, w hi ch co nver t s 2/3 of t e 0 0 5 mo les of H2 PO4 to HPO 4 2-) and d ilu te the re sulting mixtu re to one lite r 28 After mixing the buffers, the solution contains 0 75 M la ctic aci d and 0 25 M so dium lact at e. T he pK a of lact ic a cid is 3 8 6 Use the Hen d ers on Hasse lb al ch equation.

H = p Ka + lo g ([CH3]) pH = 3 86 + log ([CH3]) 3 86 + log (0 2 5/0 7 5) 5 3 8 6 1 2 What is the o n e cti on bet we en t he rmodyna mi cs an d life? 44 T he l ow e r in g of e nt ro p y n e ed . d to g ive li fe to o rg an e lle s is at the ent ro py i n the s u rr ound ings, thu s i nc reas i ng th e en tr o py of th e Uni ver Se as whole 45 Compar tmenta l iZat ion in Org an e lle S b ri n gs Com po nents of rea ct ions into proxi mity wi th one another. The energy Ch.

nge of th e reaction

iS. 0 t a ffect ed, b ut th e availa bi li ty 0 f comp 0 nents al

1. W Sint 0 p r oceed m ore r e adi ly 4 6 D NA wou ld h ave higher entropy th th e St rand s se par a t ed. Th ere a re two s ingle st ΔT - i ls te

a. Of one d oubl est ra n d a nd th es in gles t ra nd have mo re i nfo r mation al mobilit y 4 7 s ee the an s wer t o exe rc is e 4 3 I t is s ti ll unli kely that a c ell S c ou ld have ar. Se n as Sharecy t o Pl a Sm, but the q uest ion of proxi mi t y of rea ct ant s is mo ret o the point. The t hal t h e energ y c ha nge of a g iv e n r eaction 48 I t w ou ld be u ni ikely that ce.

1. 0f t he k ind we kno w but hav e evo lve d m a s s an t. The lac k of So lid s and liqui ds on which a re a t e s could form wo uld make a large d iffe

r

ene 49 The a vail abl e mate ri a ls differ from those th

t. 0 U l d have been f ou n d on earth an C onditions of te mp e r a ture and pr essu re a re Very diffe rent 5.

I. ars the u se of co nd it ions, Ore li k e those 0 n ear. h 5 1 A n um be r o f en er geti ca l ly favo rab l e int era ct io n s d r ive th e pr o c es s of Pr o te i n fol d ing ul t i mat e ly i ncreasing the entr o py o f t he lu ni 5 2 Ph ot o Sy nth esi s is end ergo nic, C r e q ui ri ng li g ht energy from the sun. The compl

all wi thout cata. ysts 14 Two of the m o st ob vi ous a d vantages are spe ed and specificity; the.

a. SOw Or kat con sta n t t emperature Or Pro du ce l it t l e r e a t 15 1 c od ing a l lo w s fo r re pr od uction of cel l S 1 6 wi th re S pe C t to cod.

n. R N A types o f n u cl eo tides have been pr o duced from b p omers in t h e abs ence of e i . her a r e ex isti n g R N A to be Copied or a n enzyme to catal yze the pr oce ss . h e ob s er v ati 0 n t hat some e x ist in g RNA m ol ec u l e s ca n c at al yze their own pro c es s i ng suggests a r o le f o r RN- i n cata lysis with this dual r ole.

N. May ha ve b een the ori gi nal in form a ti on a nd c r om o le c u le i n the

g. Δn Δi 1 Δ7 ti S Δu key t hat c e l s ou ld ha v e a r i. e nas b are cyt o pl asm wit h out a plasma m embr an e. T he pr e sen ce of the mem b ra n e prote cts c e llular compon en ts from th e envi ronme.

ta. d p re vent S th e m f r om d i ffusing awa y from e a ch o ther. The m e cell esw - thin

ac. 11 c an re ac t m ore easily i f th ey a re c lo se r t o each o th er 1 Δ 4 H o w do prokar yotes and eukaryotes differ in level s of 1 8 F iv e . i ff Δ e re nces betw een p r oka r yo tes and e ukaryo te s are (1) P roka r

yo. es d o not have Δ e - 1 d fined nu cl eus, b ut eukar y. tes h ave a nuc le us mark ed o ff fr om t h e r est o f the cell Δ b.

OU. lemembr anes (2) P ro k

ry. tes have o nly a pla sm a (cell) m em brane; eu Δ k ar yote h av. an extens ive in t e rnal memb ran e syste Δ m - 2 H. ar y otic c ells co n tai n membra ne bounded Δ O rga nel le S). hi le prok ar yo tic cells do no t (4) Euk Δ a r yo - i c e ll.

ar. Nor m al ly l a rger th an th os e of Prokary ot es, (5) Proka ry 0 tes are si n gle c e lled organ i S ms, whi l e e.

ka. Yot es c a n be mu lticel lul a r as well a s si ngle - ce lle d . P rot e ins in the s is take s pla ce on r ibo s om e s in both prokary o tes and e ukar y o t es. I n e uka ryo t es, ri bos o me s may b e bound to the end o pl a s mic ret ic ulu m or- O un d free in the c y top l a s m ; in prokar y otes, ri b os o me s ar e on l y f ound free in the cytopl a sm 1 9 W hat a re the main s tructural fea tures of prokaryot c Cells? 20 I t i s un l i k e ly that mito ch on dri- wou ld be fo und in bac teria. These euk ary o tic org a nelles are e nc l o sed b y a doub le membrane, and bact eria do not have a ll i nterna l m e mbrane s y s tem. The mito ch ondria f ound in eukaryotic c e ll s are ab o- t th e same size as mo s t b 1 6 What are the main s tructural feat ure s of euka ryoti c cells? 21 See section 1 6 for.

th fUnctio ns of the Part

SO. a n an i m a l ce l l, w hi c h ar e s ho wn i n Fi gure 1 10 (b). 2 see S ect i on 1 6 f o r th e fun ct i ons o f t he par ts o f.

Pl. n t c e l l, w hic h are s ho wn in Fig ure 1 10 (b) 23 In green plants ph ot o s y nthe s is t a k es place in th e m embrane sy s t em of chl or opl asts, w hi ch are l ar ge mem. ra ne enc lo sed organe l le S In photosy nthe tic b a cteria, th e re a re ext ens.

On. Of t he pl asma m em b ran e int o t h e int eri or of th e Ce ll c a ll ed chroma to Ph ores which ar e t he si te s of Ph o to.

yn. hes i s 2 4 Nuclei, mi tochond ri a, and ch l 0 r op l as ts are bou n de d b y a dou b le 2 5 Nu cl ei, mi t o.

h0. d ri a, a nd chl oro p l ast s a l l co n t ai n DNA. The DNA f o u n d in m it o ch ondr ia and chl oropla sts di f fers f ro m that in th e nuc le u s 2 6 M it o ch on dria ca rry out a h i gh perc enta g e of t he ox i dat i on.

en r gy - re l ea si ng rea cti O n S of the ce ll. They a re the Pri mary si tes of ATP 2 7 Th e Gol g i a p par a t us i s in v- l v e d in bin din g ca rbohydrat e

S t O r o t e i S a n d i n e X O

rti n g s u bs ta nce S f ro m t h e c e l

Ly s os omes C o nt a i n y

ro. Y ti Cen zymes, per ox iso mes Conta i n c a t a l a se (n e ed e d f or the m e ta b o l i s m of per Oxides), and gly o x y s o me s contain enzym.

S need ed b y p lants for t he gl y o x y l at e cycle. A ll t hese o r - an e l les ha v

et. e ap p e a ran c e of fla t te n ed sacs b ou nde d b y a si n gl e me m b r ane 1 7 H ow do w e class ify organ is m S: Five k in g do m s o r- ree 2 8 Mo ne ra inc lu de ba cteria c .

1 g. E co li, An acy a n o bac te ri a. I Protista. Includ e E ugle na, vo lvox, Am o eb a, and Para m eciu m Fun gi i nclude m old s and mush r oo ms. I Plant a e i nclude c l u b mosses and oa k trees. A n i mali ai n cludes pi de rs, s alm on , r attl e snakes, r o bins, a nd dogs 2 9 Th e ki ng d om Mone- r a

co. S is t s of Prok ary ote s. Th e o ther. our k ingdoms co nsi st o f euka ry otes, 3 0 The kin gdom Mo ner a is d i vi ded in to t he d omai ns, U bac teri- a and Ar chae a, on the b as is o f b i o c h e mi c a l d iffer e nc es. The d oma i n Eu kar ya consist s of the four ki n gd o ms o f eu kar y o ti c organ.

SM. 1 1 8 I s The r ecom m ON G r o u n d for a l i ce ll s? 3 1 T he major a d vant a e i s t ha t or h. v i n g com part ments or ga nelles, with spe c i a l ized fu n ctions and thu.

di. iSi on of lab o r. Another adva nta ge is that c ells Can be much l ar g er wi th o ut s ur fac e a rea t o v o lume co ns ider ati on s being critical. The ad use of comp a rtm ental i 3 2 see th e d isc u ss i on o f t h e e n d o s y m b i otic theo ry i n s e ction 1 8 1 3 3 See Exercise 3 2 The di V.

Si. n of l ab or in ce ll s gi ve r.

Se. Of greater e fficiency a n d a l a r ger n u mbe r o f i n d i v i duals. This i n t u r n al l ows mo r e opp o rt u nity for ev o lution and Sp e c i ation 1 9

HO. DOC e i s Use en e rgy 3 4 Proce ss es t h at r e l ea s e e ne r gy a re favo re d 1, 10 W ha t is the c onnection betw een en.

rg. an d c hange? 3 5 Th e ter m s pontaneou s mea ns en er g etica l ly f avo re d ; it does n ot n ece s sari l y mean fa st 1, 11 What is the crite r ion for s pont a ne i ty in b iochemica l 3 6 The system is then on Po l ar so lu te a nd water, whi ch become m or e dis order e d w he n a s o l ution is f orm ed; s y s i s p osi t iv e but co mp ar a ti v ely sm all The s u r r is nega tive and co m pa rati v e ly l arge because it i s a r eflection o f the un fav o rab le en t h al p y change for fo rming the solution (H 3 7 Proce ss es (a) and (b) a re s pontaneous, wh ereas Processes (c) and (d) are.

0t. The s pon tan eous proces ses r ep re Sent an inc rease

IE„XPe rieflc ifl saSharppHc ta pgel c©T heroi fltinatit rat, OfC ur V„
Wh„ IE theKkdd edaqidonbaSee¤ Uhistlea mOunt ΔO™-fe„0 lj
Sina llypresE n tl(f) Theprof-r·yOrapOrecu ΔI©-ats IE
adinSo1 ubieifwate Ifjed# s Whi c hinturnh-S·0 tineSress
Δb™ferinfa··ac.t Yt
hd„ thethirl-lluqeeou ·ffrshave the ΔS™-Hb„i IC e tlf
Yllnave the Sdnereld-iVeamOuhts ΔO™, eac.d dn cinS
eformi4-l·tWouldhemO IEeffec tyetogtartΔi™-e.E PEshS
el „ou Wantta Hffera tAP hasovet hepKWhichfle ans,ha tthe bas„
fromwilpredo minatewhen youth averinis ted PEMlin„itU itis-
asi ertoCO fivEft some Ofthera sEformoth eaciclnO fn tha mos„
Of„ hEacid fromplothe base ·-Ormi4O1 nabulfΔf™w·t,t hE
PHA bOvE e-hepKa't Δh©-ef.IrMp
reaomi matshmi S„Ou lobuseful ls Δa™-er.0 fa
r

efrieSofthelndivi dualAminol-ids? 2OPFOLinfistch nicallYn ot
tha minOacid„l Glycifle„ Ontainsno Ch,r„
IC„Tbo fanO fSI3 l Anamfba Ci diMwhiCh theKerUp Contain Sthero lno
wing :Anr dr oM group CSelinOtlfe„ hine YtyrOS ifleva sUlfura to
lk cysltei ne;mEt hiOnine aSe Condchifau cal r bOna tomc isO1
EUcin E th reOgni IE>l nalinOgroUPOlysinEan„
Ili„ egrOuK asP afaline ;glut aminE afl acidsfou rds ra rtate>glu
tamateina ro m atiCri ng CPhenyl a lanin E t Yrosin e·tr„ Prophra
nzahran cheasidfchainc leuc ine·valine>·l Inther„

PtdE;v dlMetuger l lieMhel ArgdsMfromlE Ub tiEdla
alll„ No acidsarEse rAr S cYsya nal tyr:theiromat icamin o acidsar„ Ph
EanthrIfa nd the suL fu rnKOnl taj nifE allinOaCi dsar EM etinaC y
s,5l Inthefep tide; GlutOTh rmal lIAS fu l lenfer mAla thrOn„
Ol„ Ta mill oacidS afeva p lI' andAla'th eacidic a lin oacidsareG, Uthal,
spIG l Aminfacidsit he rmha nthesIha l 2o arOmuhE dr>yll difica tio
norfoneOrthfco mmOnfalmi No acidsl Se erisurE„
I5„ Ofthestruc mtu rfsO-someflO di fi edalfli noacid S| HYd roxylTOl
ifleandlvdfox mflYs ineareforulhin CO lhafe'st hyr oxineisfo uhin-
hyros nOhUl inO 7i rr„
Si„e> trypto P tan,an dtheird erivativ„ Sl 8l A fllfpO-mi neOX ida-
eisanE nzmfethatd egr dedscOll nOupd sWith a nimirfar OuP i IC„ Ui in
snell ro tran sm "tte rs:C>n s efluenti Y>t hEycalfcolfur OulPers„fl
Sm Cntgal sta te9l T hE hig hconfcentra ti onofthP toPhdlinlik Pf
OtEinmgy mi di vElevatE thEleVEls orsEr olthin ;whichIEla X„
St„Ebrain lol mEtfyptop hAtin mi lkmi Sh tmflk & Ous l epyMhE rea
StatyrdIm ineincreesSholl dreMouu„h l l Thdmi„n Ol cidsth yroxi
neaflnhydr„xy prol-fleOc curinVer YfE WprOteifls! thEgEn etic-
d

edoeSno ti fIcIudet

hE„SOthEvaIE Produced ymO dification Orfty roS
nA pdProlinE>rtSp Ec t i vE]Yl3 03 flAmin oACids„ aVESP Ec i fic-
Cide Ba sEr operEie s ?83]H HistidifIE: imi caZ oleisdE PrOtongat
ed>mam, fl Oasroupis Pr e dOmilh anty dEr otOrated l ASpa·a gi fleid
minO gfOlfis de PrOtOn ated „r yptophan:ma ni flOgroupis PrEt omi nafli
y dEPO tOflated l hrO iinfam fli flOgr Ou pisPrEd OminanE ty yaepro tOni teaPlen o
liChyd roxyl„ S approxiMatE ly a 5O1 5O mixtuf OrfPrOtOnat fdalnd dEP
rotona, Ed lmfj isoelEctfic fO i ntufOrsiUtamiCa Ci d3 l25 SEr ifl
e5l 7sh isti di ne;7l 5 & lysine>9l 75 tyfOs ifle>5l 65 ia rgi fli„E >lO-
75]ldCls teinewil lhdVE nonetchal rseatpH sl 02=CU 71 + 8l 38>/2„

S„E·i tratio ncul r·eiflOl>17 mmP H=Sl 86K ml 48 >·H*3l 382 9lAf
terminxing theBuffersolUti onc1O0 mDCon tains 0 l25 M„ acti cadida-
dol 7·MOdiu flha cetat Oth enadorfa Cti dAc idisSl 86l USethE HendE
fs onHlgasselhal CHeq ult„
O„PH pka+ lo sor dl3m ·-3mPH=3 l 86l„

E„ObiCOlicIati ono fsiucOseisexerEin, cAdais·SOurc cOfenefgyform,
NforganismsSi nCl udinsHumdngs! tWu, dherEuson- hueto cfrecthe-
W
O„ Tocc sgSs tot akE-ldceclif refert mbi forderfop rO vidEPergfor th e
eAuEfg oniCon„ I Ch apted2An swelfs2l 2P hAtiS dHYd fo genBMde 1l P
rOtei nsaflamuC lei cAcidsgh dEln„
d„ Osensonas asA phiMportan tbar tforEir·StruC tulf esl o hePlicA
tionO„ DNAgnaistfa„ Sc li Pti0 fltoRN Arec uiresly dfosEnon ding
ofcomplEMEntary bas eSt o thE nAtim pA test randBy ThEc-H„
O„disno tsu frAic iEfluF rOmfro fs rEa tj yu-eAuli dist ri bMtion OfEn-
Ctro fsa titstwO Cfusl„ Isothefair cnfunShare dpairSo fEAct ronsto
sErVe asnYdr EenfondAc CEptorsfol Manymolec„ ulEsCan fOrmhY dfOEn
hOflusl ExaMple SmigSh tbEflo'CH3„
H„NH2 SfrO rato fat ose Cll„ dfA hyd fOg„ nbOnd ri t flus thaA Yd„ Yaf ose
fCO„ dnE ntl YbOh dEd toO YN„

r„I-hishYrO sen thenf ormSa lv dfosEn>Onall i tiamot he r O>MorfAol ill
a-yerOsen„ boflueldim efO-afEctica cidth e "OHfort ionfor thEC anJX
yisfU Po rmo rE C-lCh iShyd„ Ose fl h son cEdfOtE"[4] portiO-orthCaf
hoXYlg roUp dmol„
CU„E>dfuvic Ev er sal7l6 illdSe=l„ andsonitol i: 8? lib itolHi5:e
Ic ialdOnsrO lfrc afbond to wa terSalltne fi nsoxYEnb ifladStotwd th-
S

OS„ hAte mll) theUni quefi tn Essofwat Er fOrfOrmin„
hY„ Fo genbon dSaEteEm, ne stiEp fOrpE rtiesofflAmfi mpO rta nfbio
molec UlEs:wate rca pAbsOacAds anAC, d gAdabaSE'gi vinfl gi ts
reatvE, satil ity in bjOnchemiC al reactionsH 12l Ifra tofSdi dfl
otdif fEfline lc„
fO„E satiVity >ther tw ouldbeIOPO l Arso nISi This wOuld„ rAs tiCA-
lyAfe fe crallr eactionSt hAtinvOlvErq uCt iofAlsfou rSCon„
ai„ i fOxYge fo rfli trose fl hAtis mostof ma3 w hacAfe AcidSanAbas
es? 13l Cc H3 3 NHCOni„ gAteacidCC H3)3N Coniu gAte baSe) H
3N+C82=COOfC co niu gAteAcid„

H3„ CH2=m Ekc Onsiu gAteBase +H3N+C82=m Ekconiu g
At„ AcidHeM2=m Ekconiu gAt eBase) m rch 2=CO OfCconiu gAtea„ id) MrCH
2=mEkCo niugate„ aSOm=CH2=co OfCconi ugateAs eSm=C82=c„ oHconi
lgateac idmus(a) THenuferi„ aconstAnt eq αβ a= to thecontEnt
fAtioppf therroUctsort hea αβis=SOciatiofdivided byt„
e

CO„ Ce ntra tiono f the undiss OCia tEdAcid form: α H+ Ml Cb) nE qu Ali
tiveorY uaN tiAtivEd—Sc ription„ fh oMluc hacid,
HA) dissoc iatestOhyd roGenion (c) Thepro Per tyofAmo leculeIta
h

as„ Otfa ponarfeg ionina„ nonp QarfE sio nc dMea to Unofac idO
rhgsOth at dMeadfat OaburfEr„

fi.fl clS10 Wth eCa rhoxy1sr0u raaS ac1ar8e dc af boly1ateC m0Oma
fld.hEa mifl sarflp ifl jtscha nsedafor mod4y4 .4 2n Thefear e ve1y rew Si dec
nd instnath a ver flnt tiOnd l sf Ou fstofofm=43i T hEre Wi lbEm
a
ny m OfR0s Si b leOnro rfla t β·O·sfecause0ff r eerotatiofllrfunc
the pefti de t0 n| 4a/ the .
eW u Uldhenop osS ibiiit Yfdis unfide trSs lipiswithinor betweEhee
Pt dech ai5>&vi nsfll creroS sib leOnror mations T er ewUludnot h eth
e ross ibi lit Yofoxi da t β·0| readCti 0hrEac i Onsinv 0i vinssU
lfhy dry Jafldai Su lf α-dEgroufSl4 9rthebi .
di ffErencew ouldeth e lQsSof SterEosP ecifi ci . Yintufe con form
ati0 n0fanyp epti deorrfrotEin T hiSn0 Uhchal ve dgSti Cconse4ul
enCesf of thekinats .
fT aCtio nsofthEfr oreinf3 0 Ares maulpetide sP hssiolQgidu y aCtiv-
?4 6i okydci nds anisoieufile atpoiti .
fB nagaetGiη Cetposi tiOfls si tstifilu

ea dth Ei ttefta steroftheOth erchap ter4AfswErs#1ho fDcf Sthe str
uctu re0f Proteins D etEminer .
ei Functi0n2l fCa >c2)s < b> (4)s(0)c1 >s(b(3)0 lw hEfffPfot e ifli
SCnatu reA> thein tE ractiOnst hatde te rmi fesc Ofd ar Y tert ary
fldanycu atE rfla rystrucq resqare .
Ve COm ebsythe β-resence 0f th eteph turinBase n tiOnsYthe primary-
tfutture fema ifls inta tl 3 rrtera α-dOmfort io ns orAprdteinf-
flotc0 ntainstr lctura ll0 ti festfata rerEp .
at dWi th ifthe prot Ein's uchasamel ix 0fb pplEat Edsheetb Utt
hreEnti ntns i on aufeatll resQ0fthES epaf ts 0fthEprQtei nar
ere peat Edfrr Offlo fleffQ ilecu letdfQ ther0 ThIS? tiEt erm mIniS
s 0me-hitfor alls „
Offl r l42what Sth eprima rYst fuctufeo epfdeins? 4i whe na prntei
nsc 0Val ent tymoqifie d'itspr imarYstr lcture is c β
Inse4l T heP ri fllarYstr UCtu fdeter in esthef ifial tnyreEli
mtnsi O na lSt ruct urgofthe prQt eifl Themod if icatio n disfUbts
thEfOld ni g α-sl4a>stflnEtaSasm .
11 idech-ifluthatc anf it inganyfel at ifel y pola fe fVi rnmefu<b >T
ryftophanfha sthe nlarge sts idecfuai nfanyof th ec Omon . Ihi noqaCid
S>afkdit tellstore qui rEdtOn . Olare nvi nEfmEn t u< C> lysilfa
fkarsi ni negtarebEth baSicami floaCiqs . EXch afgins Ofl,f0 .t
he the rwQuidno tal f ffect the Si d edgainpKb>ifas is gnif i cantlavlSi
mi lar reaS 0n ifka pplie St Othesubsti tuti 0nfram0fl0 .
ri 01 euc in e rofall Omp 0 ldrieuci nel6 i G ly Cine i sfrEqu efl tuya
ofnserveArtsidu eb e Causeit S Sidec ha nisSos fla u aSt 0fi tiflsfal c

Estnat Wilpnot CC0m mou tEi afg eron eSi7 W he naiaining sr
eplaced byiSOieu Cine 'th erEi sn 0tEnOlg hr QomiftheE
ativeconfOrfati onf0rth ellar ge rsidec hei nof t>e i5oieufile
lOnfSequEn tuy>ther ei s asreatEnfus .
Ch nggafut le c onfOrfflltu ti ofOrthE Pro tEi ntxtatitiOse sA Ctivity wEng
lycinei sS Ihs tiftUtedint umfOrfS .leUaffe the efes en c0rthe sma
ners-dfchai fl eadstore SumPti 0n ofthEactiy Eco nfo rmatiOfl .
l Mhfors is tsuffse y0fanimAlpfOt einsa-dfat0 The tEm rerra tur e
s9 8 19 thEc o niuAlteacid .
bA epairwill actl sabff fer int hEPhfa ng eu ugl Theyhly efl pertchar .
eatpHextfemtS>a nat hemfle Cuiest end torEre reacfOthenfl w
henthefOut cuarcia .Sei5 zer0thEa fli pfac i dsca nasgr .
gA eAOfEeas ily l2 0 rheion iCais sOciat i0 nfeactio nsOfth ef oiiow .
nga min dAdas aspar ticacid v Ali ne>hi stidine>Serin e'ala-
ysin E 21 i th epaAfOrtheion .
zA ioMfrt he thi oisroup Ofc'sti fle i S83 3> s0t hisafimo aCidcou ld
ser Vasabuffr „ Ifint hEn sha ndals fo rmsoVer t hEp Hrafl Se7 33
bl 3s1 rtfl amifbgr o uPsofasfalfsin Eilnd rySin th aVe pkavalue s0
f8l 8dAlas195 .
fe Pec tiVe ly ; t hestarea ISOF0 Sib leburffer S'but tthe Ylreb othErt .
Ee dlffutheifu ffer fr ange s l2 0 AtPHs the fCa rfOx yl gfo Upisdepfo
t0 rattdat oa rSrsoxYra te>thesi deCidi ncgrn OxYiissti . Im ofEt-
alfso Pfoto fa tEd y andOfdti rO srfoup SarEp fotofat EdAtPH 7? b
otht rEma rnoXvigfOurpand thesi deCidi nc gfi-oXY ist ouPdfe depro

Ofl tEdro aca rfoxy ate'and

25 PHE=3| 8G+<0 4 8 P HE=4! 34 3 0ltuS euEhEndErson ma sse nhanche
Quati0 nfP HEPfaHo O(CH 3'm) 03 0mPHE=4 0 76+0gfc HBll D 0E=3lPHE=4 2
6+10gcol 5 M 0 l 1 0MPHE=4 0 6+0 l 40 PHE=11 63 1 0 Yes>iti Confectca
lculate theflolaralloun t$0fthE tiofor .
Sa di nserti ntotheHe nder's Ofmha sseib alchequatiOni 2 0 2g= 0l 0167
n0 ran dsl25g=0 0333m D 32 .

Thes0lu ti Onisabu

ff rlbeca lls eit Contaill Seqlla uc On Cefitral tiOnS
0 . TRIS ifthe aCid andfEe alli nefOmpS0 nhEnthet w0 S0lu tiOn ts a fEm i le-
> the-0 Cefitratio ns0 fthEresu ltinkso ll tiOnfinte- e-bSerce ofrEa
cti o Bartoio-M C-alpndor l MrRI . becallseofdi lu tiOf Th EHCl reactSwi
th haIfthetR Ispre Sent >di-i-Ingoi05 MRISCprOt On ate dr0
rnd anlor 0S MTRISC ffe eamin Cfor mp 33l AnyHlfferEntiat ha Seq-
a rqnce nEnt tiOnfOrthe acidandbasi Cdrm SWi l haveEdheq alt oits-
KaT ere f0rethebuffEr ffomex . Icise 32 Will haVelpHO f8l 3l 34 0 F
irstcalcu late themoles0fburffEr thatyo unaVe 1 00mL=0 1li .
e S>anb 1 1 iter sofolim TRIS burffEr Sol ol mOles Si flc Enthalf ref
isatitSPK a' thefelfEcqUa lc ofCentf . tiOnsOfth fdci dapplas icf°r
mpSOthEam oll ntOfRnSi sO) Oo-fll- les>afluth ea mfun t of-RSHi So-
0)5motes , fVOlthe fldd 3mlofj MH c1 >0lWi l heading OlOO3 flonesof
H+ thisWi lIre act assHOwl : TRIS+H+ un . i nyOuru fOut orsofeth
ingsWhich wi llethe H+inct i Stheiflitingr-d-efluthete Wallo uhtsc-
n

b Calcu late ea Ss hownhetoW: m+=d005 m oles+ol 0 os mOles: l oO .

mO,eS TR Is =0i 0-5moles Mi oo3m Ores=oloO 2m ole S-Owplug

h Sc Values intothEHe fde rsonma sEth archeqUation : P HEs13+l0- C T-
IS]/C TRISH +1) 3 ;9 3+l0gOi Oo-/ l Oos) PHE7 1 70 3f firstcalcllat

t emOles Of burffEr fEft ha t0u have: cw eAr goin Af0rOsfm EfOlmins offb i
ooIL =o0 1 litEfs> a fu Olliters ofpi M, RIsbuff EriSOi ol mOles Is in
cEthebu fe fi sa tpH7 7o> wesAli nex .
I ise25tha thea fbou ntOfIR isiSo0 i o2 fO1 es>and th eanQunt0 f TM-
H+isO i o8mOle s uify .. Uth enadi3m h 0f 1MH Cb yo u Willfea dainpSh 0o-
mole So fHH thisw i lIfea ctas sh0 Tn: TRIS +H+-=H+unti , Yoll fUn outo-
s0me th ilS>w hiChwill le the TRISSi . Ceitis t hElimitin fi reagen t
Al i thenIi siSOnve rfEctoTRIS HH+ =d0 l mole S TRIS=Wmol .
S hqVeus EdIP the buffEr Caqac ity ofthETRIS W Elowal-
eolo . 3flo IesOfHH+inIP pfOXi matEly0 1 1 li tEfs Ofs0l uti On T
hiSis a pfOXifla telyo .
3 H+l pHE= Al =S O 1 s0HEol 52 3 6l DH+=[Al fo mP ufleacid ' th lSKa-
H+l2 =2-a-[HAl=2 0g . H+1 :pKall 0g[A]p HE=' = {K a mol sgl^ .
} 7l u Se t heEnde rso fIna SEthaich eQuati On ofacEta tei Ofl Adc
etica cidE= 3/ 13 8 aS ubsta fice wi thapK a0 fa t 9 lasaBu ffer ran-
eOf2 1 9t .. 4l 9 1 twill notbdffere ff Ect ifE ivat p H7 l 5 l 9 u Slt hefl
enEr S0 plhasse lb alcheau-t· 0fl thErat i oofa WOu ndEs98 t0ll fo
l nAl l clses tfues . i tab lEb ufferra nsecoversalHer dn sgeof Pga+/-1 p
Hunits l a Ctica Gi dcP K3 l 86 xAlti ts Soli ulfsalt pHl 86 l 86 Ac
etiCac id cPK a= 4 76>a nd i tsSOdiu .

Sa b Pbsl7 6m 76T lusC Se tabi e3l 4 ' ka:8 l 3> inits Sprut ofiate cl
form alldit srfeea main eromm >P H] 3m 3HE Pe S Cste Tables l 4 PKa=7
9 ssinits Zwitterionic alfdi tsalfi0n icfO rm >P Hgl 5 .
l 5 4l 0 Severalfuort nehurrfr sWoul db eSu it ableFamfly TE StrEhEs >M
orS>andPI . ES> however tclEb eSt buffrerW Oll dbeMP sal si tsPka o
f7 i zi sC l best0fte esirEdfio .
7l l 42 hrhiSOlutiOnj Scdhieicd lBoom b iarsEventh olEh the Conf enfur alt-
io nofflEither thEr eeasEnor theCon ilgate .

Ci i Sol Os Ol My whyiSol 0 5o·· M-hEcorrect Concentra ti Ontoreof
t? Unre fCo nfiden tr ation slfetypi calfr 0orteate-b tes
lmof tiEtWoionic =43l A ttheeQuiv ale nfeoint0fthet-ta ti o-
aSmi ll affl untonfiCe ti Ci dffla in sise CausEofthefq lli i insi lICH
3CO 0l-HH+ofOfl the-e-sasmall vb lt nonzfO yamoufl .
Of ceti -dCidi 4 4 Bufrerin scara City isshSedyPo-te amoufhiso fthe-
Ciqanhas e fonts Pr esEntinthebur fers olu ti ofl AS0lu ti onwit .
ah Shb Ufre rningc aP acity Cane aC t wi thlarse Am0Un tofaCeta Ciobfig
Sewit h o urdristiCha ngESinpH AS0lu tio Wi thal 0fb Ufer in escap-
city cl nEactwithOnly-0-pa rati ve l ysmh l-a mn
O Unt sorcid 0rtase le fr esh owin sch aflses inth Th enmfrConCen tra-
edthebu fer rthehigsrEsi tsb ffe ri nscapa City T .
ef rstullfrert is tEdhfehas 10times-Es burffer in Capaci tiwaIthiseco
nd 'se ti ng givi nMeCm , bopaPart idaou bi b-Oldlar ac ter,
so Inpl-lca s es> theyie l iSo195 nfronforfSidu Es >tha tfeanS 60 %Yi
eld >forfore-i d l-Sts0 anufor Orts idle S'o 6% Ir hSeafefot Sa

tis fac tOryYie l-S, E-Z Ype Specific .
tY ets arou nd the prblue m0 thet WOrEp ti deSci ffEr in affli nCc
idse fu enc e burnOti nC ofPOm s0n 32) Thetitrati oncu rve sOft hetw-
p
ep. ic Swi l lgV-thesam egeneral Sh aPer therKdva l ues Ofthe la. inO
anda ca rbo Xyls r0 u PSli ni di ffEfi Ve rYcarbOxy lfo rms-wil sh Ssi-
gshfd if,e ncEsinsi dec hain pK a Wl westEcall . bof thediff er elitai
stances tot hEchar ge d sr0Us a tcHEni sof the re Pti de l suchchange .
dIf Partic ula rmMarked inPfO tE i nsl 3 9 ASP L eu mPhe tE U As-
phep hEls rhLells Pmphed L edueuh h tA SP3 4 DLF lDFD lD F LLFD-
Lp3 5 lYOUW uldg et 20 mL .
27 10 0 fOlecules 'w h ichisab 0ut Osae ar thVO u rCesl thesame cli
Cura ti o nfOrapent aPPtid eSivesmO Ec ompfehens ibiE rt's . l
tesl 3 6 l Th ey arepe i atiV e lyst a bieeCause theya r ezwi tte
riongl the,ty P ic dlyMve hi g mfEtingsP oifits 37 .
Wi, hVe ryiittle d0 U-t/ nq Compa reprta ic tinsthepro Pertiesof Ta t
Enfr Omthise (hva r0 ssetandoxfGen yine i thCra tomiCrm .
Ie Una fro-m l f ounCelltepfo Pe rti esOfthEp rotei ty ollmight beal-
letdo there Ve r seOsomfex tefft, 3 8rth es etwopEp tidesdi nfe rthe-

Ca. IVl Th eo fenthain sAs arreedl tempi nfl ldpaNerfi na l ybll tt nCyc
liCrEntid enSonly . EPt ide 3 0 tO thtEdlte rmin ang th e Nte
rmin aIsofth eoPEpC hainPe pti de can bECh arfe ddtaoPPr Opriate
PkVdiuf s> whiC . isnotth-C-SgEwiththeCyCliCPe Pti dEfmis can Pr ov
ideapls is forSen fatiOn 4 ICafb gly drat-S-renotadsou rceof th
Efi r Or enp tedfarOr biOs ymfltheSis0 filminoa Ci ds l 4 Slg 8st thaty-

cel lrai trav es ofle n hfo ma ine most obin Athi sha lt it yles> th- rti sie ssbxy sfne> dn, tteCon cen tTati On of the deo xy for flbrt heab nformaun-fl glo oj hinc, easesl ilssoaly Sncan he lo ulfdtaUSi n sthe ehS eNe dbre a thinsdi, ficultie-l ,2· Infetal hem, slobi lbtiES usu ni tconfisi ti Onis 22 wi, hr ep ulce lhe fftfth e■ chinsby tchaC hail- l thesiCkl e lce l lmuatiofarf ectsth Gich ain ,Sothe retU sfpofo, ysoUls- OfH bsha Sro ma l re ta inffll Osjobirfs , hereiati Vbxyg ln afri fi tieSiil Ow oxyel-Ore, ake pbythere tal lCeil Sr fTmthemater flalfh l sfl B eca Usfne opie wi thsickle ■celd ise aSe affchr oniC aplVa,, effli c>so flecetlswit hfetalHsir ep ro duce dtohel polerco fet hdnfa ird oxy e,, cle li Vers ystem) 55l Thecry sta lii n eror mthan seaneca, sfoxy enfn te fed Intr the qver slip> tr ans rofmi ngeoxy h,

MO,,lOtsin tcoolY he flO gtobinfl chapter 5 ANSWER s 5o1 HOWDOW e Extr ac- llrepro teins fro flct l us> l l bi fluer >P Ortte, ■E Ule sh effl> Soni Cantor2 l l fyo Untfeded to fla i n tai n thest ructU rfli fit e gr i t yo f th es UC e ll ul a r, rga n, l, es> ap O, te r ■el Ve j,.

efll,,Olldrhe le ttfl bec au seitis mre sentl elth e tis sll e.Su,, hasti Ver> m Us the SO ft en oushtolg eWi th th is de V.C e l sl sa l t ingouti sarfO cess wher e tsyani shi yiofli (Sa l tis lSe d tor- duc, hesO, Ubi l, t yOra Pr O, te i nul ti i ti tOm Gsoltofs,, lUt i o nalfuC an b e e ndfi r use d l thesal, r fflfsio ndi p Ore lon as wi th th- Iat e rin tse sOlution> which e ave s lesswat e ra Vadi h b le tOhYd,,

at, the Protcefi N ofl Ol ar lsid-C a- ■ Isbesg in ftti n te ra cti se tteen rfot ti nflo le Curlg'an d tiefy re,, Ol ti nfol ubi el ■ T he iram noacifon tenta nlatfra n■emen.

Sm,, kesomep-0-ei nsmOrts Ou Ubl th a no thers! A, fOt e i nwi t hmore nishly Pol tra mi n Oaci d Ont,, Esumac e-i-l le mores O lu blethnonf wi th- Ofeny el' O ph oicofe, ofn te leSlfnte> i hofllse ni ,,

et, e iV erc e ri sysinga, o tter■ E Vesh emhofse fiz en Sfi nte hm Og enf eat s0ox Sto s e dilfth uth un f Oke nce lsafd,, Ucie il (enfr i rus- the Sl pe rng tin ta us > oo0Xg ra nacol lfc the Pel e tv ll,,

iC,,COn tai ns t he mitfCh o ndrial d N O, rer ox i sol fs afld mi toc hold, iand l ve ovefra, finssed n, m a ■ lifCha r,, Cte rs Stics lo the rtec hi ques, Such assuc lose gfa d-e ntcllfer if ucation > wfUn dhV et obe used to, s era fa te the fty oorsnh U lSi 1■■■ ifVo Ve ainC ook in Sle atafe u, Ua l, y, lOr e than e nou sh tOd en a murt l e Proteinart Of the meat 9, PFlln di se lse Sta Ve been ■ linke dt O the ei flu n-S yst ■ l l nt,

Sb, l i ev e d th at the r ri on Pr O te in S t ravel in tel y

MP,,SYs te ■ lbl un dt O i ym P ro CY te S and e ve n t u al ly ar fiv Ca t the he ri Ou,, t iss U ewh er e the y- b esin t or ans r of l e h em f m a l Pi Onp r O te in n toth, a bo fl mat ol le o i whil e ther gar ets,

r O,, gge ne ti C pr e ds Posi ti on s t Ca Cfu l t Sc ra pi ■, hat a' ■ few i l mtc ause the cis tas er med iSe as emis t be st lr te dh Vi fise,, ti ■ laP ri on tha ta l re ad y las te a ure f eac oll for mat i ol ■ ,, P Sc l l l T- e pr O te in e ffi cie ■ c y fa ti o is ara n nit fa r y ■ e as ur,, ■ n t of the Sss e n ti ala fi na cia l con fe n t o - fas i ven ty re o fr or t o fi h 2 l E ss s ha v et h ch ig h s t PER,

13, The a mi n o ac id s th at m q st he con Sume d in the d i et b c a use the b O dy c- n n ot sy nt he Size the li n s ll ffi ci en t q U a n ti t i e s l 4 0 Rea Son s fo r t re at in gen e t i c d i s y m o d i ri e d r O s i n c l u d i ,,

C

re,,Si fle the i r pr o te in cO n te n t > i n cr e as i n g s the irs hel fi f e > i- cre- Sin st d i, r e Si s t an ■ e t O ins ect s l pr o th er pes ts > an d a ec re as i ng s th en e d e r O r P es t i c i de s t r O ,,

th,,Ml 4 l s what ust h es e d on da ty s tr U ct u- e o e Pr o te i n s? 1 5 l s h a re- O u n hi l i ty > an d t y re o f hi o l o g i c a l f un c ti o ,, C s t a ti c s t f un c t u r an er s l s s d yn a l i c > c at a l y t i c l O l O t , h e a l g i e s l) f the am i de ■ an e sas they f ■ lt e about th e an Can) p y n e a l e a r e b O the fine da z er ol l hen the t wo P l an e S w U ld f o ver l a p ri n g s u ,,

ht, at the ar b o ny l gr u p o f o l e c o n tact s t he ■ l b r t l e o th en t l A■ bu s t i s a C Om m on m o n e re t i t i v e i n r, , S u a r i t y f f U n d a n i ■ l t i Pa r ■ a u l el l S h e e t s l A l i s a i i g ■ ■ l e n t O, c u r s h s t U e e n s t l n d s f o f t h e s h e e t

causi n g s o fe si d e t o b O f O u t wa l l s 18 l Ar e v e r s e t u l n i s ar e g l ,,

n O, a p O l y fe P t i d e w h e l e t h e d i f e C t i O n Ch an ge s b y a b O u t l 8 O p l T he r e ar e t w O k i n d s - t h O Se th at c on t ain P f O l i n e an d t h O Se th at d O n O t U Se t h i s r e l 6 f O r t ,,

am,,les s 19 l The ah eli x i s n O t f u l l y e x t en d e d 'an d i t s h y d r O g en b O n d s

re,,a n a l u e m O th P r O te i n fi b e r ,, h e■ pl e a t e as h e ets t h e C t i n r i s l fo- t f U l , Ye x t e n d e d > al d , t s h , d r O g en b O n d s ar e P er f e n t i C U l ar , O t h e p r O t ei n- i b e r s l o l the aq u e O U s , t h e y a l b l U n i t o the , ■ e a ll d e r , t h e r e C e k- y , the an d 2 l l Th e s e O n e t r y O f th e P r O t O l i f e r t S i d U C i s s U ch t h at t h e s h O l d t r i t i n t O t h e a ■ i n y t- U t i l t O e s r e i t ,,

Xa, t l y r- O r a r e v e r s e t U r n l O S e e Fi g U r e ■ 2 4 g r C i l l e i

St, e O n t y r e s i d U e s m a l l e n ol g h t O r i t a t C r U C i d l P O r i n t s i f t h e C O l a g e n t r u f i P l e h e r i ■ , l 2 3 l t h e P r i n f i P a l C O ■ P O n e n t O f t O O t i s t h e P r O t e i- k e r a t i n f l h i s c h i s a C l a s s i C e X a m P l e O f t h e h i C h l S t r U C t U r e , l t h e- r i n C i P a l C O m P O n e n t O r s i , k i s t h e P r O t e i n f i b r i l l h i C h i s a C i a s s i C e X a m P l e O f t h e s t r U C t U r e l Th e s e ■ a t e ,,

en, i S s o m e w h a t O f a f a l l a c i o U s n O t i o n t h at f i b r O U s P r O t e i n s a r e a l w a y s U n i m P O r t a n t f O r t h e f U n C t i O n i n g O f l i v i n g t i s s U e s , f O r e x a m P l e , t h e C O l l a g e n s O f t e n f O r m t h e b U l k O f t h e e x t r a C e l l U l a r m a t r i x i n t h e s k i n , t e n d O n s , a n d l i g a m e n t s , w h e r e t h e y a r e a r r a n g e d i n t h i C k b U n d l e s t h at a r e e x t r e m e l y s t r O n g i n t e n s i O n ,,

ht, at the ar b o ny l gr u P o f o l e c o n tact s t he ■ l b r t l e o th en t l A■ bu s t i s a C Om m on m o n e re t i t i v e i n r, , S u a r i t y f f U n d a n i ■ l t i Pa r ■ a u l el l S h e e t s l A l i s a i i g ■ ■ l e n t O, c u r s h s t U e e n s t l n d s f o f t h e s h e e t

causi n g s o fe si d e t o b O f O u t wa l l s 18 l Ar e v e r s e t u l n i s ar e g l ,,

no, a p O l y fe P t i d e w h e l e t h e d i f e C t i O n Ch an ge s b y a b O u t l 8 O p l T he r e ar e t w O k i n d s - t h O Se th at c on t ain P f O l i n e an d t h O Se th at d O n O t U Se t h i s r e l 6 f O r t ,,

am,,les s 19 l The ah eli x i s n O t f u l l y e x t en d e d 'an d i t s h y d r O g en b O n d s

re,,a n a l u e m O th P r O te i n fi b e r ,, h e■ pl e a t e as h e ets t h e C t i n r i s l fo- t f U l , Ye x t e n d e d > al d , t s h , d r O g en b O n d s ar e P er f e n t i C U l ar , O t h e p r O t ei n- i b e r s l o l the aq u e O U s , t h e y a l b l U n i t o the , ■ e a ll d e r , t h e r e C e k- y , the an d 2 l l Th e s e O n e t r y O f th e P r O t O l i f e r t S i d U C i s s U ch t h at t h e s h O l d t r i t i n t O t h e a ■ i n y t- U t i l t O e s r e i t ,,

Xa, t l y r- O r a r e v e r s e t U r n l O S e e Fi g U r e ■ 2 4 g r C i l l e i

St, e O n t y r e s i d U e s m a l l e n ol g h t O r i t a t C r U C i d l P O r i n t s i f t h e C O l a g e n t r u f i P l e h e r i ■ , l 2 3 l t h e P r i n f i P a l C O ■ P O n e n t O f t O O t i s t h e P r O t e i- k e r a t i n f l h i s c h i s a C l a s s i C e X a m P l e O f t h e h i C h l S t r U C t U r e , l t h e- r i n C i P a l C O m P O n e n t O r s i , k i s t h e P r O t e i n f i b r i l l h i C h i s a C i a s s i C e X a m P l e O f t h e s t r U C t U r e l Th e s e ■ a t e ,,

en, i S s o m e w h a t O f a f a l l a c i o U s n O t i o n t h at f i b r O U s P r O t e i n s a r e a l w a y s U n i m P O r t a n t f O r t h e f U n C t i O n i n g O f l i v i n g t i s s U e s , f O r e x a m P l e , t h e C O l l a g e n s O f t e n f O r m t h e b U l k O f t h e e x t r a C e l l U l a r m a t r i x i n t h e s k i n , t e n d O n s , a n d l i g a m e n t s , w h e r e t h e y a r e a r r a n g e d i n t h i C k b U n d l e s t h at a r e e x t r e m e l y s t r O n g i n t e n s i O n ,,

Sb, l i ev e d th at the r ri on Pr O te in S t ravel in tel y

MP,,SYs te ■ lbl un dt O i ym P ro CY te S and e ve n t u al ly ar fiv Ca t the he ri Ou,, t iss U ewh er e the y- b esin t or ans r of l e h em f m a l Pi Onp r O te in n toth, a bo fl mat ol le o i whil e ther gar ets,

r O,, gge ne ti C pr e ds Posi ti on s t Ca Cfu l t Sc ra pi ■, hat a' ■ few i l mtc ause the cis tas er med iSe as emis t be st lr te dh Vi fise,, ti ■ laP ri on tha ta l re ad y las te a ure f eac oll for mat i ol ■ ,, P Sc l l l T- e pr O te in e ffi cie ■ c y fa ti o is ara n nit fa r y ■ e as ur,, ■ n t of the Sss e n ti ala fi na cia l con fe n t o - fas i ven ty re o fr or t o fi h 2 l E ss s ha v et h ch ig h s t PER,

13, The a mi n o ac id s th at m q st he con Sume d in the d i et b c a use the b O dy c- n n ot sy n t he Size the li n s ll ffi ci en t q U a n ti t i e s l 4 0 Rea Son s fo r t re at in gen e t i c d i s y m o d i ri e d r O s i n c l u d i ,,

C

UP,,ar e pr O to n at e d l At P h O ,, h O th e ■ Cl

Δr ■■ x- lg, O Up a n d t ,,, Si e Cr ai n d l r

bOX y lg r O

Δu ■■ r e d, P fl O t o f ■ ■ d t o a C a r ■ O

xY l at e, on e O f t h e a m i n o g r O u P s i s P r i m a r i l

Δy ■■ r- O t, ■ ■ t e l ■ a- d, h e O th e r ■ ■

il,,gro un s a m i X t ul r e O f u e P f O t O g a t d a n d e P r O t, n a t e d r O m sl 3 l t h e P l r e f e r s t O t h e r O m i l w h i c h, O th c a n O x y l s a r O l l f S a r e a e P r- t h n a t e d a n d a r O t h a m i g r O u P s P f O t O n a t e, d a t P h O 9 o 2 3 l t h e P r e f e r,

St,, t h e r e f O r m i n h i c h s O t h e r O X y g r O u P s, a r e d e P r O t O n at e d a t P h O 9 6 l 2 4 0 At P H l t h e c h, r g e d, r O u P s a r e t h e N u e r i f i f a l N H s f O r m a d u n g a n d t h e P r O t O n at e d g l a m i d i ,,

O

gr O u P O n a r s e i n i r e, s i v i n g z e r O n e t e c h a r g e l

ΔT ■■ c- ar e d g r O u P

sat P H l a r Δe ■■ s a l e a s P H i l

ith ΔT ■■ a- d, t i O f l O f t

h e c a, r O X y l a t e s r O l P O n t h e d e t e r m i n a l l e y c i n t P s i v a i n g a f e r t C h a r

e0,,■ l, 2 5 l B O th P P t i d e s, Ph e ■ g l u s e f ■ l Me t a n d V a l n r f h e c y s ■ l e,, ? h a v e a c i a r s t O f l l a t P H s e ca u s e O f t h e P r O t O n at e d N ■ t e r m i n a l a m i n O g r O u P s l n a q u e O U s s O l u t i O n s ,, Ma l a m i n g r O u P l A t h P r t h e,,

Pt, d e O n t e r i Δg ■■ h as n O n e t C h a r g e O n a u s e O f t h e P r O t O n at e d N ■ t e r m i n a l a m i n O g r O u P s ,, Gr O u n d t h e i r O n i z e d t e r m i n a l C a r b O X y l a r e Δn e g l t i v e ch ar g Δe ■■ r

eP, P t i de O th e ■ e f f e C t i v e s an e t C h a r g e of m i n a t e r m i n a l a m i n O a c i d s e O f t h e s i d e, Chain ■ l ■ b O x y l a t e g r O u P O n t h e g l u t a m i n e Δi n ad d i t i O n t O t h e t e r m i n a l a m i n O g r O u P s, 2 6 l O th e r s i d e ch ai n s O f c e r t a i n a m i n O a c i d s, s u c h a s l y s i n e, h i s t i d i n e, a n d a r g i n i n e, c a n a l s O b e P r O t O n at e d O r d e P r O t O n at e d, d e P e n d i n g O n t h e P H O f t h e s O l u t i O n ,,

ecause O f t h e si d ec

ha, na mi n O g r O u P, c b) s e r i n e b e c a U s e O f t h e l

a, kO f as i d e C h ai n c ar b O X y l i c O H i s t i di ff b l e c au s e O f t h e P r e s e n C e O f a t r a t ab l e g r O u P i n t h e s i d e C h a i n s (d a s P a r t a n d g l u t a C i d e) ,, e d r, b O X y l a r O u P i n t h e s i de C h ai n ■ l 27 l G l y C i n e i s ff e r e n t f r O m t h e O t h e r 1 9 a m i n O a c i d s b e C a U s e i t l a C k s a n y s i d e C h ai n b e y O n d a s i n g l e h y d r O g e n a t O m ,,

d, St h e r e a s i s O f a b U f f e r i n t h e b l O d r a n g e n e a r t h e P K O f i t s c ar b O X y l g r O U P ,, Th e s e f un c t i O n s a r e P e r f O r m e d b y h i s t i d i n e r e s i d u e s ■ l 2 6 l a n d 3 t O ■) ■ 4 ■ , a t t h e i r n a t i v e P H O f ■ 7 ,, Th e i m i d a z O l e r i n g O f h i s t i d i n e h as a P K a O f ■ 6 ,, a t t h i s P H , h i s t i d i n e r e s i d u e s a r e a b l e t O a C t a s e i t h e r a n a C i d O r a b as e ,, Th e i m i d a z O l e r i n g O f h i s t i d i n e h as a P K a O f ■ 6 ,, a t t h i s P H , h i s t i d i n e r e s i d u e s a r e a b l e t O a C t a s e i t h e r a n a C i d O r a b as e ,,

O, t h i s b u t e- O - h e P r a P h a r a r a l l s e m e n t b y s u b s t i t u t i n g a n O n P O l a r a m i n O a C i d w i t h a n a r O m a t i C r e s i d u e s u C h a s t r y P t O P h a n O r t y r O s i n e ,, ■ a y a l s O i n C r e a s e t h e s t a b i l i t y O f a n a l P h a h e l i x ,, s i n C e t h e r i n g s O f t h e s e a r O m a t i C r e s i d u e s C a n f O r m h y d r O g e n b O n d s w i t h t h e P e P t i d e b a C k b O n e ,, Th e f O u r t h s e C O n d a r y s t r u C t u r e l e v e l i s t h e t e r t i a r y s t r u C t u r e O f a P r O t e i n ,, Th i s l e v e l r e f e r s t O t h e O v e r a l l t h r e e- d i m e n s i O n a l s h a P e O f a P r O t e i n ,, Th e t e r t i a r y s t r u C t u r e O f a P r O t e i n i s d e t e r m i n e d b y t h e i n t e r a C t i O n s b e t w e e n t h e v a r i O u s s i d e C h ai n s (r a d i C a l s) O f t h e a m i n O a C i d r e s i d u e s ,,

V, l O r s e f l u e n c e h O m O l O g Y i s m a r g i n a l f O r u s O f C O m P a r a ■ t i v e m O d e l i ■ ■ i s t h e s t t e O t r y ,,

h, t f l e t h O d y b U t t h e n t O c O m P l r e t h e r e s u l t s w i t h t h O s e O b t a i n e d f r ■ l t h e f O l d r e C O r d i , i O n ■ 3 6) s e t h e P O t i a l t a r g e t f O r t h e t a r g e t a n g l e 3 7 l A P r i O n i s a P O t e n t i a l i n h i b i t O r O f P r O t e i n a s e s ,, i n C l u d i n g t h O s e f O u n d i n H I V ,, a n d h a s b e e n s h O w n t O i n h i b i t t h e g r O w t h O f H I V i n C e l l C u l t u r e ,,

ec, ause S, and l i ce versa, a s O l t h e chang e O f a h as t i d i n e t O a s e r i n e ,,

t h e l O s s O f a h i s t i d i n e m a y e l i m i n a t e a h y d r O g e n b O n d t h at h e l P s t O s t a b i l i z e a C e r t a i n c O n f O r m a t i O n O f t h e P r O t e i n ,, h a, C O u l d h a v e l i t t l e O r n O e f f e C t O n t h e P r O t e i n ' s f u n C t i O n ,, i f t h e r e a r e n O O t h e r i n t e r a C t i O n s i n v O l v i n g t h e s i d e C h ai n O f h i s t i d i n e ,, h O w e v e r , i f t h e h i s t i d i n e r e s i d u e i s i n v O l v e d i n a h y d r O g e n b O n d i n g n e t w O r k , t h e l O s s O f t h i s i n t e r a C t i O n C O u l d d i s r u P t t h e n e t w O r k a n d d e s t a b i l i z e t h e P r O t e i n ,,

ec, ase S, and l i ce versa, a s O l t h e chang e O f a h as t i d i n e t O a s e r i n e ,,

t h e l O s s O f a h i s t i d i n e m a y e l i m i n a t e a h y d r O g e n b O n d t h at h e l P s t O s t a b i l i z e a C e r t a i n c O n f O r m a t i O n O f t h e P r O t e i n ,, h a, C O u l d h a v e l i t t l e O r n O e f f e C t O n t h e P r O t e i n ' s f u n C t i O n ,, i f t h e r e a r e n O O t h e r i n t e r a C t i O n s i n v O l v i n g t h e s i d e C h ai n O f h i s t i d i n e ,, h O w e v e r , i f t h e h i s t i d i n e r e s i d u e i s i n v O l v e d i n a h y d r O g e n b O n d i n g n e t w O r k , t h e l O s s O f t h i s i n t e r a C t i O n C O u l d d i s r u P t t h e n e t w O r k a n d d e s t a b i l i z e t h e P r O t e i n ,,

th e ■ e m O s t O b i ■ P O r t a n t a t h i n g s a l t i t y l e s > t h- r t i s t e s s b x y s f n > d n, t h e C O n c e n t r a t i O n O f t h e d e O x y f O r ■ l b r t h e t h O n f O r m a t i O n - ■ l g l O g h i n C , e a s e s l i s s O l y S n c a n h e l O ■ l d t a l S i n s t h e h l S e ■ e b r e a t h i n s d i , f i C U l t i e - l , 2 · I n f e t a l h e m , s l O b i n t h r S u s u ni t C O n f l S i t i O n i s 2 2 w i , h r e P l l C e l h e ■ l t f t h e e ■ c h i n s b y t h e h C h a i l- l t h e s i C k l e ■ l C e l m u t a t i O n a f f e C t s t h e C h a i n > s O t h e r e t U s P O , y s O l l s- O f H b i n a S r O m a l r e t a i n ■ ■ l O g l O b i n f s O , h e r e i a t i V b x y g e n a f r i t i e s h i l O w O x y e ■ - O r e , a k e P b y t h e r e t a l C e ■ l S r f T m t h e m a t e r f a l h P l s ■ l B e c a U s e O P r e ■ i C l e s w i t h f e t a l H s i r e P r O d U C e d t O h e l P O l e r C O n f e t h ■ ■ a i r t d O x y e ,, c l e l i V e r y s y s t e ■) 5 5 l Th e r y s t a l i i n e r O r m t h a n s e a n e C a , s f O x y e n t e f e d l n t r t h e q v e r s l i P > t r a n s r O f m i n g e O x y h ,

MO,,lOtsin tcoolY he flO gtobinfl chapter 5 ANSWER s 5o1 HOWDOW e Extr ac- llrepro teins fro flct l us> l l bi fluer >P Ortte, ■E Ule sh effl> Soni Cantor2 l l fyo Untfeded to fla i n tai n thest ructU rfli fit e gr i t yo f th es UC e ll ul a r, rga n, l, es> ap O, te r ■el Ve j,.

efll,,Olldrhe le ttfl bec au seitis mre sentl elth e tis sll e.Su,, hasti Ver> m Us the SO ft en oushtolg eWi th th is de V.C e l sl sa l t ingouti sarfO cess wher e tsyani shi yiofli (Sa l tis lSe d tor- duc, hesO, Ubi l, t yOra Pr O, te i nul ti i ti tOm Gsoltofs,, lUt i o nalfuC an b e e ndfi r use d l thesal, r fflfsio ndi p Ore lon as wi th th- Iat e rin tse sOlution > which e ave s lesswat e ra Vadi h b le tOhYd,,

at, the Protcefi N ofl Ol ar lsid-C a- ■ Isbesg in ftti n te ra cti se tteen rfot ti nflo le Curlg'an d tiefy re,, Ol ti nfol ubi el ■ T he iram noacifon tenta nlatfra n■emen.

Sm,, kesomep-0-ei nsmOrts Ou Ubl th a no thers! A, fOt e i nwi t hmore nishly Pol tra mi n Oaci d Ont,, Esumac e-i-l le mores O lu blethnonf wi th- Ofeny el' O ph oicofe, ofn te leSlfnte> i hofllse ni ,,

et, e iV erc e ri sysinga, o tter■ E Vesh emhofse fiz en Sfi nte hm Og enf eat s0ox Sto s e dilfth uth un f Oke nce lsafd,, Ucie il (enfr i rus- the Sl pe rng tin ta us > oo0Xg ra nacol lfc the Pel e tv ll,,

iC,,COn tai ns t he mitfCh o ndrial d N O, rer ox i sol fs afld mi toc hold, iand l ve ovefra, finssed n, m a ■ lifCha r,, Cte rs Stics lo the rtec hi ques, Such assuc lose gfa d-e ntcllfer if ucation > wfUn dhV et obe used to, s era fa te the fty oorsnh U lSi 1■■■ ifVo Ve ainC ook in Sle atafe u, Ua l, y, lOr e than e nou sh tOd en a murt l e Proteinart Of the meat 9, PFlln di se lse Sta Ve been ■ linke dt O the ei flu n-S yst ■ l l nt,

Sb, l i ev e d th at the r ri on Pr O te in S t ravel in tel y

MP,,SYs te ■ lbl un dt O i ym P ro CY te S and e ve n t u al ly ar fiv Ca t the he ri Ou,, t iss U ewh er e the y- b esin t or ans r of l e h em f m a l Pi Onp r O te in n toth, a bo fl mat ol le o i whil e ther gar ets,

r O,, gge ne ti C pr e ds Posi ti on s t Ca Cfu l t Sc ra pi ■, hat a' ■ few i l mtc ause the cis tas er med iSe as emis t be st lr te dh Vi fise,, ti ■ laP ri on tha ta l re ad y las te a ure f eac oll for mat i ol ■ ,, P Sc l l l T- e pr O te in e ffi cie ■ c y fa ti o is ara n nit fa r y ■ e as ur,, ■ n t of the Sss e n ti ala fi na cia l con fe n t o - fas i ven ty re o fr or t o fi h 2 l E ss s ha v et h ch ig h s t PER,

13, The a mi n o ac id s th at m q st he con Sume d in the d i et b c a use the b O dy c- n n ot sy n t he Size the li n s ll ffi ci en t q U a n ti t i e s l 4 0 Rea Son s fo r t re at in gen e t i c d i s y m o d i ri e d r O s i n c l u d i ,,

C

th e ■ e m O s t O b i ■ P O r t a n t a t h i n g s a l t i t y l e s > t h- r t i s t e s s b x y s f n > d n, t h e C O n c e n t r a t i O n O f t h e d e O x y f O r ■ l b r t h e t h O n f O r m a t i O n - ■ l g l O g h i n C , e a s e s l i s s O l y S n c a n h e l O ■ l d t a l S i n s t h e h l S e ■ e b r e a t h i n s d i , f i C U l t i e - l , 2 · I n f e t a l h e m , s l O b i n t h r S u s u ni t C O n f l S i t i O n i s 2 2 w i , h r e P l l C e l h e ■ l t f t h e e ■ c h i n s b y t h e C h a i l- l t h e s i C k l e ■ l C e l m u t a t i O n a f f e C t s t h e C h a i n > s O t h e r e t U s P O , y s O l l s- O f H b i n a S r O m a l r e t a i n ■ ■ l O g l O b i n f s O , h e r e i a t i V b x y g e n a f r i t i e s h i l O w O x y e ■ - O r e , a k e P b y t h e r e t a l C e ■ l S r f T m t h e m a t e r f a l h P l s ■ l B e c a U s e O P r e ■ i C l e s w i t h f e t a l H s i r e P r O d U C e d t O h e l P O l e r C O n f e t h ■ ■ a i r t d O x y e ,, c l e l i V e r y s y s t e ■) 5 5 l Th e r y s t a l i i n e r O r m t h a n s e a n e C a , s f O x y e n t e f e d l n t r t h e q v e r s l i P > t r a n s r O f m i n g e O x y h ,

MO,,lOtsin tcoolY he flO gtobinfl chapter 5 ANSWER s 5o1 HOWDOW e Extr ac- llrepro teins fro flct l us> l l bi fluer >P Ortte, ■E Ule sh effl> Soni Cantor2 l l fyo Untfeded to fla i n tai n thest ructU rfli fit e gr i t yo f th es UC e ll ul a r, rga n, l, es> ap O, te r ■el Ve j,.

efll,,Olldrhe le ttfl bec au seitis mre sentl elth e tis sll e.Su,, hasti Ver> m Us the SO ft en oushtolg eWi th th is de V.C e l sl sa l t ingouti sarfO cess wher e tsyani shi yiofli (Sa l tis lSe d tor- duc, hesO, Ubi l, t yOra Pr O, te i nul ti i ti tOm Gsoltofs,, lUt i o nalfuC an b e e ndfi r use d l thesal, r fflfsio ndi p Ore lon as wi th th- Iat e rin tse sOlution > which e ave s lesswat e ra Vadi h b le tOhYd,,

at, the Protcefi N ofl Ol ar lsid-C a- ■ Isbesg in ftti n te ra cti se tteen rfot ti nflo le Curlg'an d tiefy re,, Ol ti nfol ubi el ■ T he iram noacifon tenta nlatfra n■emen.

Sm,, kesomep-0-ei nsmOrts Ou Ubl th a no thers! A, fOt e i nwi t hmore nishly Pol tra mi n Oaci d Ont,, Esumac e-i-l le mores O lu blethnonf wi th- Ofeny el' O ph oicofe, ofn te leSlfnte> i hofllse ni ,,

et, e iV erc e ri sysinga, o tter■ E Vesh emhofse fiz en Sfi nte hm Og enf eat s0ox Sto s e dilfth uth un f Oke nce lsafd,, Ucie il (enfr i rus- the Sl pe rng tin ta us > oo0Xg ra nacol lfc the Pel e tv ll,,

iC,,COn tai ns t he mitfCh o ndrial d N O, rer ox i sol fs afld mi toc hold, iand l ve ovefra, finssed n, m a ■ lifCha r,, Cte rs Stics lo the rtec hi ques, Such assuc lose gfa d-e ntcllfer if ucation > wfUn dhV et obe used to, s era fa te the fty oorsnh U lSi 1■■■ ifVo Ve ainC ook in Sle atafe u, Ua l, y, lOr e than e nou sh tOd en a murt l e Proteinart Of the meat 9, PFlln di se lse Sta Ve been ■ linke dt O the ei flu n-S yst ■ l l nt,

Sb, l i ev e d th at the r ri on Pr O te in S t ravel in tel y

MP,,SYs te ■ lbl un dt O i ym P ro CY te S and e ve n t u al ly ar fiv Ca t the he ri Ou,, t iss U ewh er e the y- b esin t or ans r of l e h em f m a l Pi Onp r O te in n toth, a bo fl mat ol le o i whil e ther gar ets,

r O,, gge ne ti C pr e ds Posi ti on s t Ca Cfu l t Sc ra pi ■, hat a' ■ few i l mtc ause the cis tas er med iSe as emis t be st lr te dh Vi fise,, ti ■ laP ri on tha ta l re ad y las te a ure f eac oll for mat i ol ■ ,, P Sc l l l T- e pr O te in e ffi cie ■ c y fa ti o is ara n nit fa r y ■ e as ur,, ■ n t of the Sss e n ti ala fi na cia l con fe n t o - fas i ven ty re o fr or t o fi h 2 l E ss s ha v et h ch ig h s t PER,

13, The a mi n o ac id s th at m q st he con Sume d in the d i et b c a use the b O dy c- n n ot sy n t he Size the li n s ll ffi ci en t q U a n ti t i e s l 4 0 Rea Son s fo r t re at in gen e t i c d i s y m o d i ri e d r O s i n c l u d i ,,

C

th e ■ e m O s t O b i ■ P O r t a n t a t h i n g s a l t i t y l e s > t h- r t i s t e s s b x y s f n > d n, t h e C O n c e n t r a t i O n O f t h e d e O x y f O r ■ l b r t h e t h O n f O r m a t i O n - ■ l g l O g h i n C , e a s e s l i s s O l y S n c a n h e l O ■ l d t a l S i n s t h e h l S e ■ e b r e a t h i n s d i , f i C U l t i e - l , 2 · I n f e t a l h e m , s l O b i n t h r S u s u ni t C O n f l S i t i O n i s 2 2 w i , h r e P l l C e l h e ■ l t f t h e e ■ c h i n s b y t h e C h a i l- l t h e s i C k l e ■ l C e l m u t a t i O n a f f e C t s t h e C h a i n > s O t h e r e t U s P O , y s O l l s- O f H b i n a S r O m a l r e t a i n ■ ■ l O g l O b i n f s O , h e r e i a t i V b x y g e n a f r i t i e s h i l O w O x y e ■ - O r e , a k e P b y t h e r e t a l C e ■ l S r f T m t h e m a t e r f a l h P l s ■ l B e c a U s e O P r e ■ i C l e s w i t h f e t a l H s i r e P r O d U C e d t O h e l P O l e r C O n f e t h ■ ■ a i r t d O x y e ,, c l e l i V e r y s y s t e ■) 5 5 l Th e r y s t a l i i n e r O r m t h a n s e a n e C a , s f O x y e n t e f e d l n t r t h e q v e r s l i P > t r a n s r O f m i n g e O x y h ,

fContaminatiRPTHEr iVa tivesthis ef forsows Wt-tlefumbe rof Cvcu Grufunt-l-Hereqifuth, tuWdmjnodcidsar ef eleased-fl- qdar-
mOuntSran dlOucou, dflottuellwicho newASSuproSed tOsethecOrre-
tOne142 untefrir stCIc1e> the firsta fiSCco fham ifloac id sfrom[heW
tE flmin aRthWoul dse reactedandpr eEasedas PTHeri.,
at,VEsHo livOUlcusEtaOubleSisHaunflain OtknOWhiclOreWaS thetr ye=ea=
=44 4l %4j9 tiaSressi bu etatyOu, PTo teinSmo tpufeana needsadditio n
alPuf ificationsstepStudfri veandSingle pOryrepticdeOfli.,
aL,,OP OSsimbletha tthe, rOteinhS-Ubunjts's Om UltiPlePoryPepti de
chainsc Olld beyi eldi ng the contradic to r'freshtsi,d Therealetwof
fasentns-h-thaveCate fmi nidha tire fo tly sifeoral.,
ifl, nE'Wh iChis WhattryPS- rigsspeci ficrorfNO,r mUuytherewOuld, eon
tyO neEragm efuTendling Withara···inla cid,t hEtwas notARSOfL,
Sanlwevou idiqme diate rykWwt hAtitw,Sth ectetrm in tpsHsti di
neisanas icorfoteinma,t thOug hitaSysua lly neuntralan at herefore
do esrOtreactUi,hat rysi rUt is pOssibletha tifit,,
eP,,efIv irOnnfntOf th Creactio-?,h-histidiffewas fo s itivelYc hareCaflu
wasrECOs,,iZe dby tryPsin4Ou t twOllderlyfo,arEgativeCoffen tfatio-
Ofth-V-riOusAmin,,dc iasthisiSim Portant hE Cau seitwO uldhEpY Ou
Pla flyOurSelle,,

Ci,geexPeri mefnthetten ForeXaMPle ?ifYOuRdaP
rO,einwOsecOffro siti Ons hOWednea rOflatic aMli-OdcidsP itwO uldeaw-
SeOftiMeto usEaCh YOtUYPsi ndigEsti on! 4sEcyan OgE lbromide
Wo uldheusetESSaS,herei snOtth iOnin,,rrYPsinvOluMen itEtes-
tEr as theProteins,,5%basicre sidUes! Tryp Sin WOuld,,hredthe
PrOteinfitOO VerOpieCES,W hiCHOuldbeVe-yhara cto aflanyZe4PCh
YOttYPsi nwOuldneasi0, choicesThe rEarefOre thanro, rre SidUesOf
arOflatiCami,,OacidsI Thei odmiPoaCid-PrOteinfOu ldsecltUrOUrtim,
St,,O SSibIyYiEld ing niCerfasIe nsinthe=3 0AmiPOaCid-
nestIbwhi Ch dPreSeIle nceEeff ectiv elYbythefa flaNdeflad ationsOoI
tOuldworKse stirfthetasi Cre sidUes Te rES prAdetOu tinthePrOte-
in
Dl,,nUnatfay,f ragmEnt singthePro persiZe rapSE WOuldhE shn-rateCo
ragIIE, uFOrtue bSicEffSiduCts wereiflhEri fOtuera,,iloacids,thEre
w OuldbeEffecting fra gIentth atCo UldnOtbeChapt,,rEANSwersdLw
hEmakesEnz YmeSSic hEef etiv e BiOlOgiCalcaraL,gs?DiEfze Sa
rEmAfvor,,ErsofIasni tuDemOreef rectiv eEas cataLYststhatar eBo-
enzYma ticcat,,
lY,,tSi 2Th eppaio fit yorEnzy Ie sAfePrOtei fs,PuffOmEC afaLyti CRN
AS (fi hOlyntS la rekrOw n 3) AbOutESeconds Iy Earxi,,VEn t/1o7 vent-
X3s6adys/ea fx24 hOus/da Y$6o OseCOnds/hOur= 3Iss Ec OnadsP,,
IE,,ZyMeshOld thESub str AtesIn fa VOr abIes rAti alPOs it io nst afnathe
ybindIE fre cti VelyUnfEt rAl nSi,,ionstate, toStAbili zeiN OfEth
Ata licatMaIy stslowE rune acTivAti onenErgy; sOth iSiS nOt ari-
ticuAl fe MMle=62 whatpISt,,
ED,,freferen Ceb etweent heskineticanMASpEcIfIcIneractiOnS ?sO Thef eaC
tiOr gnucStEith oXYgeSfStherfOu, natiCaLlyfAVOr ed 1s@7l If th
eprOteinwerec Yt osoliCtonc, tEMEtuswerE hr Oke nore rY vOlcdIude
nutrifuEat1OO,OOOKa,s Inadi thEOrganc IleswOuldchEinfIfredIet,
YO,,rdZynewO uncteinfthESuPErNa tant talonswith allthEoth eficy tosOli,
OfIsIn sO natUrE mitOchOndria Vi adifferentia, OfsucrO sesridiE
ntcentrifugationO use afOther,,OmogEni zationfEchniqueo thin
ewithdistrngEe,
erEnft tcort leasetheE nzymef rOfth emeIfInefO TablesE Xi stO tEi
ryOlhOll manYgrdmsO rdlmon iUmsUi fatE cas >tOadUrOctacertain, Er
cenfusaturatiO n AsOd prIanmO ulDe to tuketh ChOmOgenzA te anUadE
nOusasStOyi,,
Id,,2OOgsatu,rAtEd sIl utiOfl IEt,tesalfl PlOSi trOrtsfll inUtesonic eanE-
hNc enUtEr irfuSer serA,,a tethe SiMmata nufrMthe PreCi pita tej AS
sayshOrtheProt einYOud rewO rki nsIit hI AdaIlbreast oth
esurem natantt,,
ar,,iveata 4OsatUratEasO1 UtiOflfur eP eUtthePrOcesSint hisfaY; Yo
UWi lfrinOO,,tWh Ilthe Pe rc Ent saturatiOn ifl ASneeasrOhetoPrEc
iPitatefiU lOrEasOnabLynar,,
hh,,MOgEniZatiO flwOu ldheabletO inCrater hE sOluUbrOtO-
einMfrom thEfEroEisOmes;WhiC harEr rEsiEn cEn tri fUAti OnduSOOO
XsOUUsgEdimEfluthEmitOc lpn driaC b fOlke nOrinra ct)hesUP-
rnatafaIlL,,UdtheNave Prote in XsOu tmProteifM Free zethawtC hfli
quesandsOnica tiOn,,
UL,,ACc OmPliShthESanE thinsOr-hUll-tOCrOpdria Ind cUtiP erOxiSOne
sCo Unthese Parated ifli tiallyNIs lEnts@grad ie ntcentrifugatiO
s,,zwAtIsC UluMchrOmat OgrAphy?1 l(bsize< bspEcifiCdisa-
db,fldi negAbiLity,<> netChargeI2 ThEi AnestPrOteifnseUlteffi
rS ttuESma-LS-EluteIagtOla rge PrOtEins areexcIUde dfr OMth
iflteriOrOfth EgelbeadsOthEHave reSSaVai,
Ab,,EQUmns PaCeto triveIlESSe fUtially? they travel,,Sh Or t erdi
staflCE afterUtEir stI13rACO lfEO unicalbEeruted byraisinstnes
arEcOfEntra tiO nofMa cdinGAmbie liSAfkatiatas.

hiJerA ff iflityrO fUheifOUnd PfOmteifl thanthE sta tiOnary
resinIisa fkdOesIS
aI,iSCheApe nsUtUles SPregCifi cAspCifiCIi Gndfla Y,,emf-
sRec ifiCbutiSOfthE xPEfls iVeI4O AcdPOlfdCaneEiu tEdby rai-
insthC,,altC On centrati On orlyChding insthEPHI sAt jSCh eAr bu
tit,,ightn Ot beaSSPeCi fiCrO fa PArticuLarAr,,tUfl chanI gingthe
PHmAYbE nO respEcifi cEO rati ghtPIranCEb utextre MESOf,,
HM,,YdisOue-A-UreEthEPro tei fO 1sO raisinfEtPEsaltC OfEncentrati oni-
rUd,t iVeUsafE MOstPrOteinSWill QlutEthi SWayafAst iUbeact i
ve inUtecIs thfanefzyMEl,,
hE,,altcdpEr emovEd la tervia di anYsisifn ecSSar yI cha ngsinsth
ePHepOuth tofemOv,,thech Ar gE EdItaSet hE PrOtE,,nStob tOfl
dentIty readdMPrO te ifls Ar rOrtsIy ubiEa tthEisOE ndtr,CPO jntsU 16
)TheqAsis OfMOstPrOtEifls isIgAarOse, cE lLUIO Se,,deXIrafl,olfO.
YO,,l7I See hiAqIres 1 7l1sNIa t hinthEfrac tiO natiOn rafEOfaSe fraltrat,
OfIc OlumnfOrEcuI esWiIe lUtewitha linEafrEta tiO nphiFOFMOgrAPhYErs
lSt hEi rEJutiOflO lumnsIASe riE sOf standardsfcamb erU ltrOs rAntiafdiz
ethECOlumfln>andthe,,
afl,,MknOwflcainEae terminEdbYfIle asurifls tS E lutEOrnOIu meAtj cOmrari
ngroastAnlara curvEj 19I fOrtPrOtEin sWOuldeI-teifthEvEi,,
VO,,lmetOGfuE rallWuIld pOtuesEp arAtEdI 2OIYESYth eIlaflYasEfOuI dc
OmeOutintheV Oid VOluEs butthESA VOlkIbGrfJUedi ntheCOlUm-
bIa canfdOuIdEIutEmOfE sIOwIy2 1usETurAn ni OnfChdrgECOlUmn
SuChSOlsE PhaNSEC Qu AtEthNafYamiIES,,,,UtthECOlumnAtPHsrSs-

Ptide cha ins<2)SidECha inMhOndS>inflUYiflA pYOssIbIEhyd rOgEn
bflduOfbr OrgaccEPtorsO nutE side chains<3)hYdrOPhObicgi
ftt,,rACtiOns;inVOlvingtue fOn polarsrO urpsonthEPrOteinx4)elec tf
OstatiCinfera,,ti Ons;in tOW inSalfychaIrse dsrOu psonthEPrOteCin
s<5)mEtalligatiO nfinflYOrinsCOrajflati OnOnds,,

e

tW,,EnsiqE CAlinsandamEtaliO nJzOds tabIiZatiO nfanrCOnfOrmati-
naEp enason to WerigO fEfErsy, regarale ssOrthEkindf. nfEra
CtiO nin volIed Wi thSqaUlbriE cuEntErACtiOns;the mainCO-
PO,,flt OrflErsYchange,s iN thAaIpICwitnFEla tiveYs mAUeflUfO
pychangEsI nthEc,,sEO rmUlecule sSiar-E-SProteins, the nuMbe-
OfpOssiblEc onfOrmations makEs theenurOpicAlrEO renerSYching
esmUchMfEi mportantO This isSpar,iCu lArlytrUEintheca SEofhYd
rOphOhiCinteractiOnsI4l sWhatstHErertiPstrUCtUrEO.
Pr,,tEins27 seE fi gure 4)2frAhydrOgEnbOnd thAti sfartOrthe
hEli XOse cHO nargy strU, tUfE)lsEEriSUrEs1 13fOrAhy drOgEnbO fEi
tha tisPart,,
ft,,rti arStrUCtur ecsidEchaiflhYd rogEnbO nding)128 lseErig Ure4
1 13for EjctrostAti ci nteractio,,SStUcha sm isghtbesEe,n tEE
nthesi dechAinSOfiYsin eanOus,,arAtEd2 9sEeFiSUrE4 13rOranexI-
PtOra,d isu lfide bOn dO 3O) seErigsUre4 13fOanExample,,

fhydfOp hO bicbOnd

SI,,1O AchArErOneisAp rOtein tuAtaiOsa MOth Er ProtEifline=O
1,inscOnfle ctyvalEdeIs itr rOfasbOci atinEswi tho the rProteifls
bEforEitha sfelchedi, Sfi nauMat UrEOrM3 2)OOnSi sur atio
flr erEnstOthE,,Osition OfgrOUpsdu etO cOv alentbOndi ngEx amP
lesj ncUdEsAnd is omerSInclOPtidIlis OlfUrsIco,,
f,,rmatiOnferdfS,0th EPOsiti On insOrgrO bsi nsP afEduet O rOte-
tiOn,,roundsinfle,bOnds lnexa flE Ie is thE differEnc Eb tWEEn hEli
P Sda nastasEr,
d,OnfOrmati onSO fEtha fe13sI riErO sSi briefe atufes,limitOssibIEfO
teifl conFigurAti OnsA nalOn,,

tO,,rmatiOnsIO > AlthOUghAny OnOf20 AminoAciDS is POSsiblEAtEach
POSitiOnOnA ,,

On,iSUsEdbASEdictatEd bythEseqUEnCEtha tfOuIs fOrthatPrO
t,,ifl <2)EitherA pOsfa flUAm ifloac idtOu rdeuSEdAtEaC hPOsit,
OfexCEptFOrglYcine> hUtO nlYL eall,,
fl,,ACids Are UsEd(3)The PE PtidEgrO uflsplanArSot hatofluYcAn differa
ngements AreOnSEnYtEr thEr ransformiSmOfEstAblIsEd,,
S,,hAsOneUsUallYfOUndinPrOteifls<4)theOrsIesAfluCtheMOSt
t
i,,allY takEonAn yvAluEfrom1 toOOO >butsOme ansIesalrEnOtP
OSSibIb,,CauSeO fsterlchAnGrAflCe:ang,,

eS,hAtara fESterificatIyAllOwed mAYnotpHEstA
b,,lizAtiOn sinterActiOns? SuchAsth Os Ein thEhEli X(5)thEPrima
rystfUctUlreActerEr,,ipesanDP tiMyMhertiary Structure> AcCOrdi-
gto thE SeCOndAtrUctUres En,,iCOde)34)rE OmiCallY,Coll
asenAdsAnuAte rnA-ystfUctUrEtyPEcalsEi tAgsAlu UtiPlEPOlY PEPtidE
qhAi n,,HOWEver,Most diSCUssiOnsOf quatErnAr YStrUCtUrE inV
OlVesubUnit SO fgIO,,UlarProtEins,notfibrOUSones likeCOl-
agEnAnYS,,iEn tistsCOnsidErthEcOllAgEntriPle,,
El,XOrEAneEAMPleOfasEcOndAry strUCtUrEinfOgEnOsidum dOdECYsu,
fA,,EPOlyAcrYlAmidEgEle ctrOPhOresisliswithSDS MEthE charGEAnd
shAPE differEnt SOfProtEinsArEmi nimIzEdsOthAtthEO nlYPA
rameterdEtErmiNingthEMig,,

OA,,AndOfCOiI Sh,P,,

IT,US;chAngEAndshArEdnEt limitEdto 3d imProlyAcrY lamidE
gE,U seaforselfiliratiOnc hromato grAPhY;thEnAme -P
rOtE inScantr,,VEr AroundthEb eadsYstEre byhAvilashOrte rP
AthfOtrAtEralEdchAininGAsterIIitEfEct rOnOre sistheProt,,
iflAr EforcEdtO fOthroughthEmitrixOSOtICy travEl mprESsiOnlYi
fthEvAr ebig fE rEct allSEt here isIll OrEfric tiOnI37 I TEMWi
S37?0 cO cAtiON 9sO 4HWD-wEbEttE rMinEthEP,,iMAr yStrUCtUre-
fAProtein3SITiCE cpptidEsfA dditiOnWI 1AVEthEidEntitYO fthE,,
It,,fliMusinitsfi fs tcyCIEs SOdO in gasEparA teExPEfim Ents isn
OtnECE ssA rY3 9I tmight tEllyOu ifthEfOrMi nwasP,,fOr ifthE-
EwerEsubuni fs l4l Thf AffUnto rgAdAnre agEntEmUstExactlYmAtc
hthEA fOun tofMEtmifliniterfirstrEActiO,,
II,the reistOrittrEAm an rEAdEnt>sOmeOft lEttermi fliAl mOtrEA-
tI IftrEAfis tOO fIchSOmEOft,,
ES,,CondamifOa ci dWiIrEa CtI mfit-erCa sE>therEwi lI leasfIn dm Ou

α-helorfectthree dimensionalstructureofthepro tein nterpre-
e
nc. Onsoundsubstratecanmakethepf-tein naruentdae rnl turelo the re-
ultsofinterv e thatthelec hl α-isisisforrectsec ausefesll-tsf, omi-
fferent eX rerifents co lucontra diC ttheproPos ed mechan is
m
I I, thatcasey tne mechpis mwOuldiave etoreflouj lrfecto.
CC. fflllo datethene wexperi lle nta nes uttsi9ith epr ese nCe orac
atalystar, eCt Stferateof dr eac ti0 th estamodr, erfeee le rgychang
ei α-alterflodynati. Profer tYt hatdoe sflotd epenflont, eractio
nrato consequ enfutl9 th eP resepf eoft Becaug lYs thasffer feeth
α-0 thepresente Orac data lysto Verstheac tivatiOllentr.
YO. a reactionru l b en zymes, like aquidatalysts, ificreds etherate
Ofth-forward anarever se rea ctionto.
hE. ame extenu l2 n th ea mOu ntofPro du ctO b ta, nedinarac tionfae-
enfdso fthueeq, in ibfi Um Cofstantu Aca tal ystdOe.
NO. affectthat16i3 howCan weDe scr ibt enzymtk, netics inmgathem
atika mathemather eactionjsfins to rder with respec. tto A/ f-
StofaE fwithrespectto bal naseQna ortErolera lb Tleadefai neaffech
ani smofthe, eactio ni slikeif tpiny Onleo nelDlecule ea chOr-
andsn d TheEas ie sudytofOi rolft, era teO-thSre actiOnistem-
moni tofthedae c reaSei · atsOn, afcedt34ofn, r efrEctint.
ed. Sappe dran cEofits v rnfusefa phnter fo Uld nttvstasoo dlaytOm
olit o nthefdte Ofethre ati on! YOu-fEPrOb ably fUmpinsthi Sre
aCtiOnifnbu ffersO.
ut. Oftokeepthph re latively constantlu f YOuartfbt runnfast, ert
action inanlifre.

S

O

Illi On y Our unt herisofac id en aturdt iolO fthe nzY me 16 E
n
zY. est enatuhave rai rnvsharpP HO ptimumra luesl tisnece, sarytO
epsuretha tthefOrth ereaCtiOnmixtu reS α-aysathEoptim fnuflydru-
e
Th. SiS es pecialf tr ufforeaCtiOfS. ha tr eaUifforP rO duCe hyarOg-fliOn
s) 6 04HO. Do sUbstrate Si in atOfnzy me s2 r0 nfl th elockanqdk-
yflod. If th esu bstrat efitsinrgOcompara, iv eiyri gi dProtin.
hatnaSanaCtiVesitetei thaWell ade fineds hapei nterindicedl.
it. Ode p tween zymeUnder goes acon fO fnati Onac.
afl. cOnbi nationst of ubs tra telT leactives itetales shape around th
esu, str atd 1819 T heESCO mplex. OUld nanatien ers ytO Usn, w
ith acomSeu eftia lfidr geaC tivati, onfre rgtoth etransi tiOfstd-
0 2 0 Aminoaci dsthdataperara, art ilfth effi nfacid seq Uencecdno eC
lose toe aCho, her int hreeCiffens iO nSbecaus eofPrOtei nrO
ldlinsfTh eCriticaldmfba.
id. affein theact iVesitu 2 nfh Overallp roteifst ructu reisnt
ededt°e fSu reth eCo rrectarra n gement of aminoga-idsinthea
ctiveSite, 22 Th es tron ginh iti tio nfinic atS tightbi fld instoth
ea ctiveSi, dT hus'tlfc ollpoundisv ery likelytoreat ransition.
ta. Ean abstle d5 whata feSome exAMPLeSof enzYMe ca tal yzedReac
tiOnS?23 I seeFigsur esd6. fldl 7124 N Oralhentz yMgsf °
°MiCh aelis tMEthenflan Qetic sh T heki fletiCb enavi0-oralio-
ter icentimes dOeSn obsCy theMiCrale.
iS. MEfitentqua tio 2 sTh esgrap hOff atea g ainls ts
upstr-tecorfen tratiOnisSismoi dgurOfan al ds ΔtC ic EnZymfbuth Vp
Cro1 icforanfzy ne thatObe ysthe IME nte nequatio.
I6. 6What isth emichael is MEnthnAPPr0 achtO E-Zym tk inetiCS?2 dT
nereCtiOfive locitYre tlainStheSalle withfl cr tasi ngenzYmecofcent
rtiOb I tisfreO. eti cantypOssibleY bu thishlyuth ikelyforfl-eaC
tiOntOtes aturgateqwithfnz yflel 2, I theSteq dY.
ta. eaSSu mpi dfist hatt fEc onc enfta tiOnfth eSu lst thatfecofl
plexfOesn Ot-hag eq p re, i absV0 Ver thet inti nW hichth exP efi-fl
enttakesP ract.
hE. ate orgappearan ceoft hecomP le Xis fetequilt oi strattor di sappea
fanc eSimPlifyMi nstheequa, t iO nsfor enzymtkin, tics 2 T Ufn
Overmuth er=Ymth 2 use equation, Ilo1 cal=0 1svmtk=cb >v=0 3sV
max=(bv=()oomax=cdV =0 67ma X= (bv=h ohV. Oseegraph, v max=()
681mY=fK M=0 42M 1Ose
Ab Pi7 I Whatdretne Ilo del s for the Behavior of all ostficdizy.
es. I ain th fc o nfterredmodei >d lthesu bunfi tsinrinal. OSter ic en
zymgar efcu fai ifnt heSant-form, ei th enth e1or the I they are i-
ndlinsfTh eCriticaldmfba.
id. af fein theact iVesitu 2 nfh Overallp roteifst ructu reisnt
ededt°e fSu reth eCo rrectarra n gement of aminoga-idsinthea
ctiveSite, 22 Th es tron ginh iti tio nfinic atS tightbi fld instoth
ea ctiveSi, dT hus'tlfc ollpoundisv ery likelytoreat ransition.
ta. Ean abstle d5 whata feSome exAMPLeSof enzYMe ca tal yzedReac
tiOnS?23 I seeFigsur esd6. fldl 7124 N Oralhentz yMgsf °
°MiCh aelis tMEthenflan Qetic sh T heki fletiCb enavi0-oralio-
ter icentimes dOeSn obsCy theMiCrale.
iS. MEfitentqua tio 2 sTh esgrap hOff atea g ainls ts
upstr-tecorfen tratiOnisSismoi dgurOfan al ds ΔtC ic EnZymfbuth Vp
Cro1 icforanfzy ne thatOb e ysthe IME nte nequatio.
I6. 6What isth emichael is MEnthnAPPr0 achtO E-Zym tk inetiCS?2 dT
nereCtiOfive locitYre tlainStheSalle withfl cr tasi ngenzYmecofcent
rtiOb I tisfreO. eti cantypOssibleY bu thishlyuth ikelyforfl-eaC
tiOntOtes aturgateqwithfnz yflel 2, I theSteq dY.
ta. eaSSu mpi dfist hatt fEc onc enfta tiOnfth eSu lst thatfecofl
plexfOesn Ot-hag eq p re, i absV0 Ver thet inti nW hichth exP efi-fl
enttakesP ract.
hE. ate orgappearan ceoft hecomP le Xis fetequilt oi strattor di sappea
fanc eSimPlifyMi nstheequa, t iO nsfor enzymtkin, tics 2 T Ufn
Overmuth er=Ymth 2 use equation, Ilo1 cal=0 1svmtk=cb >v=0 3sV
max=(bv=()oomax=cdV =0 67ma X= (bv=h ohV. Oseegraph, v max=()
681mY=fK M=0 42M 1Ose
Ab Pi7 I Whatdretne Ilo del s for the Behavior of all ostficdizy.
es. I ain th fc o nfterredmodei >d lthesu bunfi tsinrinal. OSter ic en
zymgar efcu fai ifnt heSant-form, ei th enth e1or the I they are i-

oteinXmiSan et negativeCha rgeP uta hom Osenate co ntainfing.
roteinx ofnt hecomfan dW ash wi thut hestart ifbUfferpro
teinxfil tsinfdto the corufflO thenenufeb yfUnningsa sautgfadien
tl 22 I Useacationexchan geCo lumn> lik eCMCP ratise >and.

Uf, tatpH 6 Pro tfinxwith a velr0
Si. ivecharge fdWi listicktO, α-eCOufln123 Wi thaqUaternary, a min
e tufcol-flnf esinfdiways.
as. Net posi tulec tri fleeyhuyou do n thiveto wor r'aboutthe pHofOurb
Uf feraltern gther orflOft hecomfO wit h'tertiary ami ne thf-
eiS dicisSociableHydrOge nsdnit hereSim.
yb. Positive ofre Ultra lychargfsed cerfn din sontheb ffe rentl2 4 Thee
asies twaYw, uldbetOUseasu. rosS Cpfdie, flnt Ose rarate thfm ito
cho nanidr rolfthe re roxiSO mesfrir stT hefln, fgaOP.

nt,emi to Cfin dria via hatsth Omogepiza tiOn rSonication)
CC. tri fdeth emitOcfiOn dridi mere ile tfOUudo gntainpr otE inBwhil
euEsure nfi atantC. fit ainedPro teifA Icon tam inflant gcOUldgst il
lexi Stvhlt the ycansCie aneCrawA ybytinni nseelri tfat io bOnSEp
hdeiGm7 5 whichHO uldSe, ar at cfrr tmAalleb' dn dtheCnitne, x change ch
Dm dOgra phfotO nsep ha fOsea tfhO q PrOteinb WOUld neneutrdl.
fld. OUi cdute, whi lepfo teinaw Ou ids tickatOtheCOil mnt2fi, lUtaflic
acid wiltEcu te dfi rstsecdUEthgCOi UfflnplisCi OseOtits.
Il, Cu ciFeandu y Sineui nlpfOsit, Ve1 y changfed aflodf in istiOcrO theco i
umf roel ute ue inc'ra ifetiEp. toarOUnd el' roEluttUy Sine-
aisetrPht ta fuhuflO 2e0 amofOr af mOnicesO Iv entiil-MoV, then-
flo ram-flonicidsrastes, sO α-hetyldanarew il Usetiefi-Stt.
el. ter, oil owed by sdicin eafld the pSutdmicacidc 27 Inefion Pdiarfam
inOacids, i il Stic kthe nostOtUestatiO fary Phases gnutafic.

Ci. Wil lmOvEthe rastefStrOn lolE dlyg1 YCin ta fndthePre fl
Yl. Idn inf2esflAp rotdin sO iutiOfferro marfammoni umSu lra tepre.
arationisEats sedOver'ase ifilt rati°
ndiumnWher'e heprotei nSO, interes t'
il. elute int hevoidVO ill mer thesai t'bei nsgler ysflai vw i ill moveth
ro UsUle colU mpslo tiVin thisfay 'the pro teinsWill eavetheselt.
Eh infdindex, ttecolu mnW, tio Utit is whatdsli ect rophorets
is?. 9isize, shape, and Char ge'sO-gar, seafn dP Ova, ryiamid-
l 31Po, yaC ryia mide 32lDN, istheE-IeC ule nOstOfne fSepara
tetona galise, ie cEff opfhore Sis, ath Ousn PfO tansc amlsOpES.

pa,a ted l 33'Thosewi ththehighestchargE
/M, SSratio woUldMO, etherastes tl T hfeder ethf eevdr iab les
tOconsid er and fo ste ifctfOPh oreseparfOnfa fiwaytOe timina.
etwoorf hf Vdfi abUessOthra, the sdmdr atiO nishy Si zeorhYcharge b
uf notbYfo thH2e.

Ine 31se egraph: vm

aX2I 5x10-4M=1, KM=11 6x10sW 32 iseqera Ph: K M=2, 8X 10M=C 0
fl eflutionsWe renotd ter Mfin ed d ifectu-l Absofbanc evaluts Ve
reusedin S. eacha samatter O, rOn vEnienceI 33 is Ee graph: vmax
=, 132x 11-3 M=, KM=1 0 2 3, 1 0-3M 3 4 the, Urflver nflUm ferisZ0 4 3P
e rmi nU. e35 I Th enflmeCrofm oleS ofepZyme iS1 I 56x10 tru mo
VernlbEr=10, 7o osec-1.
6. The10WKM fortheam mat icdlinflac idsindicatestha tth eMichaeo
XidizedPr, ferEtia lyO 37 Ut iseasier tOdetectVeiat ionsOfind iV
idu alpoi nts fOmaStdai sshetlinet, alfnroma cufVetsu theasSump-
iontha ttheMisafli nitiatio, ofth Ebindin fsaffi lity beEfluEnCes-
ist rateCanat, CEnzmei svaUidWE nth erat eOfdi SsOCi ationof th
eeffZymE subsUbstrateComplex.
O. fo dU tandfizym, isSmuchsma lle rtuant lera tEo fdissoCiati ofl ort
hecomPl ex tOe nzYMEand Suss tfate1 o7 howOftenz 'in ati cReAc, iOn
sresfOnsi DI-hibitors?3 9 I nthECa seofcomPetit -veihi bitiob t.
evaluEofKMEWhi lEthe
value O-KMre m a in-unch, nfEdinOnc dlretiti veinhi
, 40lAc-mpc e-ive i nhib-tOr-bi OckSbindin
gnotCatal YsisI41 A
no nCompe itive,
fhinitiora cE SroUna fE t eafr initOfrth
enzYmE-O r tssU bs tratefOrmi
it
Orbinfst Ot heIctivesi t
eofa nenzYm, > Pt ve ntings
bindafsOrthesubstrat
qAno ncompetitive inhibi-or-bifflsatsi tE diffEr entro
mthe-c i VESite, cluin gaOinformatiOnal
C. anSe, Whic hrEflectsthaC tiVesite issOble tObinJsuh, tfatmuic onv
ertitOproduct, 43 Icompeti tive inhibiti onafle ove rc
OmeJVad dingEnOughsubstrate b.
t his is Ot- ruefOrmi-Of m
SofenZy me inhib
t. Oll44 ALincVE aVer Burk PlotisusEfu lnEcaus eitsives astrdish thi-
e Utiseas ier tOete minEpOlWe1 lPOinsfi, tOastrdig hUlfeth an
tOa curv0 45 in aLi, nEweav efi Bu rkPlOtEfO fcompetit iVein hibi tiOn
, the lin eSintersec tfatc the yaxisin terCepE Wh iCh isEqUa ltOl /Vmax
I I ma BurkPlOtfOrmin competi ti ve inhibitio n, the lines
as. hOWnbY thefle gati vefre eEren Ecfha fhe1 Th efactma tsrucO secanfEmd
i. tai nEdinaro Ysergtmo sPhEr tisarEfi ecti ofOrthE kinEticas Pe ctsof-
hef eacti, Ib requiri nsoler cOmin sgdnaC ti Vati ofthEnef sytarr ie n of F
Is tEqUe s timOnO Ster.
b. bly ; locallcon cen tfl tiOscm ds sdc tion Cincepts> Cou ldeasily di
ctatethedi, ec tionsecOn, qUes tio nsPOnsE ab ityfotiloca tcOnce
ntra tiOnsfOUnd seiEdfhes.
f. Di ff-ttQVEl
C. Me dre lati veUla rgE cl o f fdi 3kcalinth Ereverfs era.
to n
see' h ow- ver, the
dOla Se fEa ction ifi rstOly si sl o7 f heat ifns aprotEin defiatu rfsi tl
Enzymfti cdct ilitYaepen.

sequa l'o n [illegible] 4
2MW s, v ma x² 1 5i - m m, l m
in-l5 nO^n c OPPe,t i t' ve, n h

ib, [illegible] Competitive inhibition Km does not affect Vmax

n*e

The rate of the maximal

esSm ew ith in the limits of re rr or some of the conditions she si ven to ne sis ific anti fig.

rel 4 s l' t- i s ve r' g Oo· i

n the case of non-competitive i

nhibitors; much of me'ta bolic o n

tr.. ld ce fers on fed b aCk in hi bition my down Stream on.

p'ti

tive in hi bit'rs i- e question i

SPer la rSm ootin the case for competitive inhibitors, which charmingly confirms Michaelis and Menten's

d z n V d s o mean ti bi o ti' s, h- W

eve r, a re' o mp ti i V

ei nhibito r s(g O, e- for these ick pe r

SO.,? ba d f or the ac te ria) las BOt hunes, ope and the ilte r ref' Swill

chamber The hippesw, lun tents ec tab O vet hexMisatn es ati ve a, u'sO

fl/rsh so n NO'a l' r Dsdrugs are en'z y m in h

ibit Or s' b u ran j Mor mic ia ss of such rugs sin hibit sth e Hi

v pro te ase, you would find ounce

r Standard ec on c'e p t So- s u b strate bi n di ng

inhi bi ti' n), a- d i gh

hit of bi nding' 5 1 l A' n i t' ve si b l' -h, bit

Of is bou ndb y co la ue nt ion ds N On co V alenti.. ter ac ti o sa

re la tively we ak and ide a si mple k el s 2 l A no nc O ffset it i v d hi

bitor do s fo thi nato phalcti ve Site or alle..

ZY..e l t Ss tr l' ture of the a mor e ation not ha to rties us tra te h..

p'r

7answers 7 l l' p o es' n e Michaeli s

l Men te n Mo de nde St ra the be ha vior of a llo steric en zy m S?

ul Al l0 ste ric en z n' Sd Sp la y si

Sm oid al ki n e ti cs we n ra' e Sa- e

P1O t l' d Ve- s U l s S

Ub st ra te co nce n tra ti o M i Cha e l

iS- m ef lue n z m' St h i ti h y r' n d uck in e ti cs A l l0..

ter i en zy m es usually ha' e mu- t

i m P l e S' b U l i t S' a na- h

Cb si nh ro f Sub S' e- fa re so re f f e c t O

rml ec lu es too ne su s ul fa te ha f fe s th e bi nd i fe b chav io r of the o' f Sub lh

its' z l Ii, d ne n zy m at S edi n the e ar l y st a' e s Or Y tidine nu cle O tid

es yn l es s l AP ac, t Sa s a pos, ti v' c' f e c to l O, ATc d e an dc tPac t Sa san m d the

te rmi Ns ho u l d u se u s e d r of en Zym e s th at l is P l a y Mi chael i S- Me fl ten ki

ne ti cs h- us i ti s fn t, Se d wi th a l l o ste r i Ce n z Y mes l' t ec hn i ca l l y Comp

e t i ti ve a d hi NO f Comp e ti ti V e in h, bi ti O n ar e a l so te r f i S th at a rter

e st f i C' e d r o mi c ke l i s l l en t e fl en z Y me S' s a l th O ugh th e f Co..

CE..tSa r e l' P li c a b l e t O an f en z Y me l A fli n h i' - i t' o r t' a t b' n

d s t o a p l l o ste r i Ce f l A Z' n g - t t, e S q m e s i - - as h e S u b s t' a t, e i s S i fi l i

ar tra d l a s s i d u c O p p e t i t i v e i n h i b i t' o r s m a k e t h e s l a p e o,

th.. Curv em or e s i g m o i d a l d l l 2 l W a S t h e s u b s t r a t e C o n c e n t r a t i o n f h a l t h e a l s o h a

l f o r t h e m i x i a l v e l o C i t y l i s t e r m i s t h e c d w i t h u b s t' r a t e c o n c e n, Y m e

s w p e r t h e t e r m K M i s n' t a p p' O P r i a t e l 3 l A m r e U r y c o n f r o l h d h a

s u s f a t o s e r a r a t e t h e s u s f i n t s o,

AT.. a S e l W h e n t h e S u b u l i t s' r e s e r a t h' o n e t' P e o r s l b s u h i t r e t a i n

c a r a t a l' t i c a c t i V i t y b u t w a s f o l o r e r a l l O s t e r i c i n h i b i t e n b y c t P r T h e o t, e r s u b l i t y r e h a d n O A c a S e a c t i v i t y' %

#5 [illegible] 82 What a fa f e th e C h e m i c a l n a t u r e s o f t h e l a P i d r y p e s

2 l I n f o t h e t y p e s o f l i f a d s' g i v e e d l i S e s t e r i f i c, t o C a r s o y l i c a c i d s ; w i t h t h r e C s u c h e s t e r l . f k a g s f, o r m l i n f i a l y, a l C f o r s a n e t

w o i n f o s f h e a, i d y l e t h a f o l d m i n e s t Th e s t r u c t u r, a l d f e r r e f l e C o m,

Si f t h e P u t u r e o f t h e t h i r a s t' r l i n k a g e O s l y c e r o l l f h o s h a

t i d y l e t h a p o l d l i n e s' t h e t h i r d f a t' o x, l g r O u r o f s l y c e r o l i s e S t e

f i f i e d f t' t e d c a r b o y l i c,

AC.. d b l l t' o P h i S P h O r i C a c i d t h e P h O S P h O r i C a c i d m o i e t y i s e S t e r i

f i e d i n t h e m t' o e t h y l a m i n e a n d c h o l i n e s' e s t' e r s a n d i s l O s. a l B O t h S

P h i N g m ye l i n s a f i d a P h O S P h a t i d y l c h O l i n e s C O n t a i n P h O S P h O r i C a c i.,

eS..e r i f i e d t o a f f i m i n C a r c i n O g e n i c W i t h c h e m i s t r y t h e C h O l i n e - t h e - e a s e o r a p h-

S P h a t i d y l c h O l i n e d f a c i d m a y - e c h o l i n e i n t o C a s b o f a s p h i n s m o n Y.

li..l Th e y c d i f f e r i n t h e s e c o n d a l C O l o i t. O f f i c h P h o s p h o r i C a

c i d i s e s t e r i, f i e d i n p h S P h a t i d y l C h O l i n e S, t h e s e c o n d a l c o h o l i s

O I.. S a l' C e r o y l W i c h i s a s a l s o f o r m e d e s t e r s b y o n d s t o t w o c a r b o x y l i c a c i d s I

n s p h i n s o m, e l i n s ? t h e s e c o n d a l c o h o l i s a n o t h e r f a m i l i a r c h o l i n e b a s e s p h i n g

o s i n e ; W h i C h i s f o r, m e d f r o m a m i d e b o n d t..

he..zYme have insach a fficte ristic fa ti O f the Dal I f the se q uen ti a l mod,

l' th e s u b u n i t s C a f f e c t i n d i, v i d u a l l y f r o m f l' h e s e q u e n t i a l

b e h a v i o r a n e x p l a i n e d a t i v e c o,

pe..a ti vi ty be ca use a Subst rate binding to the fro m f o u, d i n d u c f O t h

er Sub l' n i t s. OS Ti t c h f o t h e r O f f t h e f b.. r d u c i n g b i n d i n g a f f i n i t y

o r s l g r e a t, e r C o o p e r a t i v i t y s t r a y' O r e a t b' y h a v i n g a h i g h e r r a t i' O

f t h e D' a r O m m i t i a l S o r a V O r d' b y d i v i n g s u b s t r a t e i n t o

he..d i S s o C i a t i On con stant for the Substrate bind ing to th

e

Tf..f l' 1, y l' m e l' a v a l u e i s t h e c h e m i c a l p o t e n t i a l d i f f e r e n C e b e t w e e n t h e

d i l u e i S t a n d a r d s t a, O r t h e d i s s o C i a t i o n C o n s t a n,.

ts. O r s u b, s t r a t e a n d t h e t r a n s f O r m a t i O n f o r m e n z y m e' s u c h t h a t c' a R V E r. i s l a n y m

o f t h e s a r e P o s s i b l e l e v e n e, e r t' a l l y, k n o w n f o r s u r f a c e' t h e n z y m e w O

r k.? f a the r w e r e d i t e a f l O d e n t h a t e X P l a i n s t h e O b s e r v e d l e v

i o n f t h i s v e r y P O s S i b l e t h e r a l n o t h e r m o d e l w o l l e c t e d o s O-

s w e l l 7 3 h O w D O e s P h O P h o r y l a' t i o n o f s P

e f i c r e s i d u e s e f f e c t u a t e' l o l Aki n a s e i s a l l e n z y m' e t h a t P h O s P h O-

Y

la..e S a P r O t e i n f i l s i n s a h i g h e n' r g y p h o s P h a t e, s u c h a s A' P a s t h e p h O

s P h a t e b o n d i n 2 O S e f i n e' t h e f e m o l e c u l e' O f t h e C h e m i c a l

p r O d u c e n e' r g y f o r t h e C o n t r a c t i n g m u S c l e s C o n t r

a c t, a n c a u s e m u s C l e P h O s P h o r y l a s e t o b e c o l l e c t e d i n C h e m i c a l

p r O d u c e n e' r g y f o r t h e C o n t r a c t i n g m u S c l e s C o n t r a c t i o n P h O s P h o r y,

a t i o n a f f e c t s t h e r e q u i r e s t i m e f o r n e c a s c a d e s e q u e n c e s. i n S w i t h s t r u c t u r a l d y n a m i c

o r d' i n e p h r i n d i f f e r e n t i a l s t e r s b e t w e e n t h e a c t i v e a n d i n a c t i v e

s u s f a t o s e r a r a t e t h e s u s f i n t s o,

ei..Pho S f O r y l a t i o n s o t h e r e s f o l s e t i m e i s S i o w e r p h o w e v e r t h e

c a s c a d e' e f f e c t p r o d u c e s m a n f o l d r e a c t i o n v a r i a t i o n s h a v e b e e n

u e s' s o t h e e f f e c t a r e a l O l a r g e r a d i u s o f a c t i o n t h e s P h o r y l a s e c a s c a d e

P h O s P h O r y l a t e d t h i s P, O S P h o r y l a t i o n a r e f l e c t i v e

m u l t i p l e f e e d b a c k l o o p s t h a t a l l o w f o r s e n s i t i v e a n d r e g u l a t e d r e s p o n s e s t o

e n v i r o n m e n t a l c h a n g e s a n d h o r m o n a l s i g n a l s. T h e s e l o o p s e n s u r e t h a t

t h e e n z y m e a c t i v i t y i s t i g h t l y r e g u l a t e d a n d a d a p t s t o t h e c e l l' s

n e e d s, m a i n t a i n i n g h o m e o s t a s i s a n d p r o p e r c e l l u l a r f u n c t i o n.

Ch..ll Our Y ns i n o g e n e s c o m p l e x S i f i c a t i o n V e f o r m i n g m u l t i p l e

Ul..d i g e s t S i t s e l f f u r t h e r t h e p r o f i l i n g s t r u c t u r e s i n a c t i v e f o r m s i t u f t s

o u t y t h.. [illegible] i n o g r o u f o f t h e C i S o l e u C i l e p r o d u c e d b y t h e r i r s t r e

a y t s e i s n e a r t h e a c t i v e s i t e o f t h e m o t y r o s i n A S i t u f t s

s a c t i v i t y l 2 0 z m o s e h a r e o f t e n S e e n w i t h a l i g e s t i v e e..

Z Y l l

e s t r a t a r b' l l d i c e d i n t e s s u f a l a t e d i n a b o t h e n

l f t h e n z y m e w e r e a c t i V e i n f l e d i a t e l y u r o n f o u c t i o n y i t w o l d a d i g e s t o t h e r

e l u P r o t e i n s. l f f e i t w o u l d a U S e g r e a t a m p l e o f B y n d V i n f i t p r o d u c

e g a s a z' m o. t h y i t c a n b e s a f e l Y m i d e n t h e f l t a n s f o r m e d u t h e d i g e S t i V

e t i s s l k e s u c h a S t h e s t o m a.

h0..S m l u i n t e s t i n e ? w h e r e i t c a f l t h e n e a c t i v a t e d 2 0 t h i s a

l l O w s f O r a m o r e r a p i d a f d S P o l s e h e n t h e b o r m o n e i s f e d e a t h e

o m p l e. S d i r e a d l s y n t u e S i Z i d d i e u S u a l l y i, s t r e q u i r e S b r e a k i n g O

n e o r t w o s h a s t o m a k e i t a c t i v e) t h e h O r m O n e C a l l e d i s e c a n d a r i l y

t o s f l l d e m a n d 7 0 s H O w D O a c t i v e s i t e e v e n t s o r a l m z' m e f f e c t t

h r e a c t i O n e s O.

Se..i n e a r d h i s t i d i n e a r e t h e t w o f o s t r i t i c a l a m i n o a C i d s i n t h e a c t i v e-

i t e O f c h y m o t r y P s i n 1 3 l O T h e n i t i a l P h a s e f t e a s s t e f i r s t P r O d..

Ct..n d i n v O l v e S a m i C y l a t e n z y m e i n t

e r..e d i a t e) T h i s s t e P i s f a s t e r t h a n t h e s e C O n d P a r t o w h e r e f a t e f C O m

b s i n t o t h e a c t i v e s i t e a n d b r e a k s t h e e n z y m e_ a c y l b O n d) . 2 l I

n t h e r i P s i s t e P O r t l e t e a C t i O n t h e s e f i r t h y d r O x y l i s t e n z y m e i n t e r m e d i a t e

t h i r d a t t a C k s t h e s l b s t r a t e f e P t i d e b O n d) i n t h e s e f o r d s t e P r a t

e r i s t h e f u c l e..

Ph..l e t a t t a c k [illegible] t h e n z y m e i n t e r m e d i a t e f r o m t h e f i r s t P e p t i d e

s l s f r e s O, s t e P s i n V O l v i n g h y d r O l y s i s f O l l o w e d b y

[illegible] n e r a t e d c a t a l y s i s f O l l o w e d b y

r o g e n a t i o n s e r i n e) s u b s t r a t e a n d e l e a v i n g g r o u P a s e l t h i s i s f O l l O..

ed..m e d i a t e l y b y a n i d e c a t a l y s i s t e P w e i t s i V e s t h e n y d

r O.. e n t O t h e a l l i d e g f O u P. f t h e P e P t i d e b.. n d t h a t i..

ea..i n g s a s i m i l a r s c h e m e t a, e s P l a c e i n t h e s e c o n d P h a s e, f t h e

r e a. t i O n m e c h a n i s m t h e f i r s t P r h s e i s f

a s e f f

Ors eve fa t a s f l s) t h e s e r i n e a t P o s i t i o n [illegible] n t e r s e c t a t t h e x

[illegible] a x i s i n t e r a c t

) W h i

Edbylin ki fingsman

ernits Cmo fmmfrs) tos&tuE n APr Oreinis PO ryMlEfrorme db Y

1. Ski fisa mi fMlc id$to&eth Er.A ₄ c l£gcaciqi SaPO lymfEfOr ₙ e dbyli nki fig n ucP0tid Es tuse ₜ heI] catiby sis i sth epmce $sth ati fcret se$the fa teofChEfi cdn reactions. Om P.rd to th Cup Cata lyzed re aCtiOn B i o l0$ic dnca tayststsa repr0t ein Si nu mbs tal ca£eS; th eon, y EXc epti olsar ea fewt yP esofRN A whiC hc afcatgaiy zesfMEo rther e a cli on Sortheir 0w fmct a1bjis m! Thesge fct iCcO cieiSt hemea psb yWhiCh theinf0 fMI tio fr0rtEts tfuctu eafdfU fc tio n orfIIiV instheing $si Sp asSed fr0m oneqen era tio fhotfmEXu theseq, gfNCe orpuri fles afldPyri midine$in DNAfa fhi- $the gEne tic codecR NAistheC0in gll teria li fsom eVir tSes >1 o2Wha tisthe df„ica na turOf fMPo rtantBiofMl fCules?zl .h Ec 0fr ec tm at chOff unCti₀ 'al ' groups, 4n dcomPOufus Contain ifng that fun ti onll-rolPis¾ive n1fth erO1l0 Winsgli Su AminogrO thPCHSCH2 NHzan$oxY1g!o„

PC„e ton e>c HgCOCH3 HzdfoWisrO

U„CH3OHCarfOxYl gr OuPCHCOOH car boxYl grou Pc aldehY-E >CHSc HgCHOthi O1sfOupc„ 3Sifs teri fnldEcHS CO OcHzcH3Dou ble bofdc HzcH CH CHSAmi-E ii nka gecH3COON.

C„3) zether dgcH2 ocfCHBSi therfUcti opalsro„ PSint heCom POun„ SfUl roX4lzer 0re1s28 ?the con CEnt ofvitaliSfhEi dth 4Organ, coll- ounfsc 0lud„

fl, Yhell-deby liv ifigS-stensga nawe resbey„ fldt herE-InO fia ioflatofvin vestig at iofSWoh fEr! -sy fthesis Showedt„ ato rsa fi cOm P Oun ds? li keino rganiCo nes?, di dn 0t Fe dlired Vit aniSti fex„

1„ Drat ion? bu tra thenfEObey eathatlws 0f chemi str Yand P hSicSan- thalsll er sth., eCtto nab 0ratory inv$Stigatio n suSsecnu Cht\ )? t hecl- CeP tw asextflae dtO them uqflorecfmPle X? but $ti ll testa b1?d.

S„ipi-eof biOchemistry 5 Urea; li kea1 or $hni cCJmroun ds?- asth esamfIo l e Cuidfs truC tur ewhet her it isproduc„d- ya livin-Org anisMbrnou 6.7 ! FiVE7i ft h£twOcf CIOPT 0P a flederi vativ esa rea1 OWf dl s? mipTe„

fl„

iff ere ntalcon 0ls?1 la lcfh Yde S/k etonES?a rnll0 Each ePo XidaESa fldet herSH10 3Wha„CEn BiOchEmistr ySSF Ab O„

t„0ss ib lerfig -nsofLife?9! lti Sgefler dl lYb eti eve dt β h- tfar bni Sthe1i kE,Yh.

S. Sf 0fa lliife f0rms,ea fthmbOund ore xtrater restridl 1 olE ighte„ hrESi cluesflo„ldgiveZ0 l8rd 6x1023 roS Sibi li ti£s! thIS 1o1es.

dUS WOut dbefle Cessar Ytol4 Veat tEa StiVosfla r o! Sntmg belK 6l

02„X1023 >Ofr0ss i, i li ti es nil Th enU MEr iS4 4 0r 1 2x 1oz 4 Wiich istW icEAVo g adr O's number 1 1 2 l .NA iSC a pa bleo fro tlcodin„ afldca ta lysi s1 a1 C atalfsi Sa how sli vifs 0rgA nism Stoc afr yout„

hE„iCa l reactiOns nil chm 0re qff-CiE ntlyt4a fWi tdutc atdl ysth 1 4l tfoofthe Dstoph Vi0 Usadvanta ges arespe tdan dSP ecificity; thEYa 1SOW ork atCO ₙ staflutefll Perfatyl reolPrOduc„

li„tieh eatu1 ₄ cldi ngβ-11OWsfOrre P roduct-OrO ₙ C ells ₁₁ 6! Wit„ resP tCt to codins R NA ll tyrepO-Ynu Ci tOt idsha vetE enpfold ULce df romm ₀ n Ome„ Sint heabSencfor ei thE rapr ₑ e XiSting,NAto heco Pi edOr ₐn enly met0at alyzet„epro C essi T heO„ SelVatio nth ats„m e-Xis ti n₈RNAmol ecEs canfate- lze thei„ Ownpr 0mEss ing Sug„eSt gurol erorN Ai rtatal y sis! wit hthiSd„

al„Ole ?RNAm aY navebe entheir ig fdi info rma ti Ongnacr om o lecule ifl tEfOrisinorlif„ l! ltiSun likel y tiatC„

llC Oui dha Vearisemsb arecy to Pla smW ith-Outap1 asmm embfa nel-thePresEfl

CE„fth em efllbafleflr-Otfct scEl ₁ β-ialConfon entsfrOm theenlir ontfdan dpre ventS, hEmfrOmii rru S in4way frOl ₈ aCh0 therl -th β- nolecuie swi thiga-lic anf eac tmorEasi-Yif„ heyalfecl0-ert oeach0 th En!0 4h owooPrikary Ot β-saldchlkary Otes Dif ₆ e rip levei sof wl s! Fived ifferetic es-tweefl-fok drio tesaflell-ary ₀ tESal e< D Prokary Ote sdn0t ialE awel1„

dE„irfan„Cle us!butfuk af yotesha vEafl Uctelu smilk Ed offfr0- the reStfth e cEl ₄wadio lbiemeflrafla (2) Pr okdr-yotesh Ve ₀ nlyaplasm aCcE lD mefl brane e lika ry0tesha vEal neNtersi VEnte mllm.

flb„ane s ystem (3)zasEU kar0 tiCE ll sCOflaint mrgne lbounded, rga ne lf-) whieProkar.

Ot„Cce l lSlonou(4)Elka l fOti ccEl ₁ Sar effOmmuyl arager th ant hose- fPr0ka ryOte-l <5)Pro kary0 teS a1E-ilsmeg-Lie aOrganas„

S?„hi ne eukar yotescafl β-emlltifcell U narisl-Ina ₛ S ingqtem ₄ Pr0 ₑ e ns y nthe„ista kEs placeonfl ₆ cf O mesiph-thP fo kar yOtesa nacU kar fote S! In ₑu ka ryOt ₑ S ? fi „Osom eSm gyrEbouflat otfEe nuoplasmj-fEti-l lUm Orr ounafrEgin th ecytOpl-flsi ₙ P r Oka fmo tS?i boflO nesIr ₀ OnlYf ounf dfree ₁ fine ctOPlasl 1 ₉5th„ tare themdinstru ct ufl 1Fea ₜ ure SOfP r Ok ₙ r YOt icce ll S ₂₂ Ol ,tiSu fhik elytha„ mito C hOndriaW„ Undberdin dfl bAC,E ral T hesteukar yotiC Or santhEkar enc nOS e d YAd ouble- elbra ne>a fu fdc terii adofor ta veaninte-fnal „emra nesyst Efll The„

it„Ile hO-dnafro unta nelka fyoti ccEl ₁ Safea hOU tth„Sam e s izeasmos- ₁₀₀6wh dare theM ₙ fns truCtu faiF eltures ofre ka rfot iCce 1s? 21 s e se ctiOn 1 1 6fO r t h.f UNC tionSor thefa rtSO ₙ an afli d 1Ce l

1?„hi Ch-reshow nitrigure h10(ab12 2lsfes ection.)1 6f o fhefuncti- sO fthep artSo fP lan tcel ₁ w hic Δ ʰ are Δ ˢ howfhri ₉ ure lho b>l - 3u fnare elPlantsPh ot OSYn ₜ eSi stakeS PlaCti fnthe mem ₉ ra nesys ₄ tEfof ch 1or OPlasts? Yh iCh are1 arge me mbran e encl„scEa or Rafel fES ! Imp rot Osyn the tica ctre nd? thEre dttX tefS ₁ onsOf thEr! as fMlle mb ₀

anlin tothE inte rioro rfhce il β-anfEcdhro lmatiPhor eS ? wiCha re thesit-Sof ₙ h4tOSYn thesiSt 2 4! Nlc1 ei/mitoC ro pori ₉ bapl-h1O ₀

OP.asts arebOll nuedsya d OUb 1@25 ! Nyfcei ₉ mi tochon dria ? ani chlor OPl astsa1 Icfh tqaflo NA! Th eDN-rounda nmit ochpl-iaa„di fCh Hro- las tSdi fers rfOmtha tin th@20 Mit OC ₙ. hdriaca ₜ ry oUt ahiSh Fefte nalg eof th eoxidatiOnEn er gyfrOdS ifsreactio nSorthe cEl lmrE ydret hepr1ma ry sitE sOfAlPn2 ! th eGO ₁ giap rar atisis inlO ryedi nhnaifng Carnohydrfates to rfo ₙ t, ifnsa ₙ d ne xpOrt in ssh- ₉t- fice

tv aCidl ₍ Pe ri dure 8l 6)O5 ! thi SuiPi di sa Cera mide, which iSo- ekinaQ0 sPhin ₉ olipid 1 6! sphin 3O1 iPidSC- rtain Imide h0 nds dOff Ote in$ Bothcdhave„

Yd,0 Phobica flaty drOphiliC

Pa„tsrandb oth CanOccurl ₁ nfce llmembran es?bUthe ir fUnCtiOnsar- diff ere nt! An ycolfbinpation offatt4Ci„ .Si Sres ibl0 sis tefbi β-scOflai flac hara cte risti C-Use„ Imi nSstr Uc ture?Wich0t he rlipi- sicflo tO wax Es dre este fSofl Of8Ch gincarnoxy licdic ₄ dSa nuOph- Cra i nhcoholsIth eyte Cfat Ors Cerou fldsPr Ote Cti„

eC„ati nssh-PhOs pholipi ds Irem orehydroph-ict„ anch one ster ol!thEP hOPhate g r Ou pisch arsed wanth e ₄ tUch ea lc oho lisCha rgedfro β-alf thesEsgOupsi nt eractre„

di„Wi th wateri Cho lesterO ind So nuyasinsaltPO l4rgroup, afl- OhUlO t hewaxEs ur racEt oatinsi sa1ar fierth ₄ threv EfltslOssofva ter13 rThesu rra Ce!ax keepsprod Uc Erfe shb Yprfeve fltuiles„

SS„fiaterl 14 ! choleste ro lisflotVerYWa ter SOlu bl e>ulti ecit„ ini sasCool natu rai Ete reatb whiC his actual ly partorl iPOP fote i nstha ttraflsPo ftfhEl„

SS„Olu bsrefltShrOlg h th eb loo d 1s! The ₁ cithi n th eess YolksS erYesas a rallU sify ifl gaserb, forl li nsec! OEave Sic lesterifi P idsint hEbutter Cfre quefltly tri acylglycerOls a fereta rfl inth eveS i Cle-anlonfo tfor maS Cerate ₁₀₀ l OT hefem Ov alorf tEo ilalso remo Vesthepatu.

al„i usinwa Fe sonthef Eatherst THE yll ustrgf-ner ate Beforetheh rasC anfe ₙ sl 3hatIsth en aturEordi Onosicalfem-ranes? 17hr. aVlaryceroi safenoI-Ouni dnalli mA Im EfbrgantSu ₙ s, tat fm EntsC-)and(dDireconsi Stentwi thW la tisklOwna ₙ ouflEmbrdne gi cOvdne ht bOnginsbetlee flli ri casAndr oteins fs tat eflent ₍) ocCUrsi nsomea fc tOr i fmotifSbut i sfotwi despfEacio th er wiSO PfOt eins ₄ ur ifith eli Pidbi lay ₍ e rsf ath ert hanbeing San dw i Chethet Weenthenf sta tefler(a) ! Bunkiemfl Ecule ₛ tEnato bOrqunin nde Ou terli fidra yerstatEme nt(b)!!191 The public.

Sa„tine d tOthEi de aorPoryUnsat urdte dfatsAshell th ₄ fUll Te ₄ mCOn figura ti OflS iveSamoFepara„

abie C OnS istEncyhrece fltw?h Ow Every Co flcer ns ha1e ari sEra bOuttheExtefl ttoflj-hsU Ch P fldelfc smmics a turate„

fats ! 20! partiall yhyar Ogenatedvese tab leoi ls-ave ₜ heaeSir

dC„is ter fyrolmanfyr ocUs Su Oasll eo flama rifE apatOmrOn, ft So rfVm 2 ! Many Ofthedub le bondsfve beefaturfdeac r ₁ sCOcOntainsfar tiar β

Yh dflO ₉ eltetv ₑ setabl ₀ ists ! 2a! The reiSl ₑ S-hfartdis ₈ β seass ₀ cidted-thd jetsIO ins atu ratedf- tty ₄ idS ! 23.rlet- ansitionte mpera ture ₁ Sto„

er, fa β-ipidbilay ₑ fWi thflOs tryursa turated fatty acidsCOmpare dtOn ₑ wi tha hig ₙ fe rCe mtaseQfrs dtu fat e dfat, dc idsT hebi ayerfwith ₜ heuns at urfa tedfatty dci dsiS„ Ife β-dyflbre ₄ d-sqrtereqt ha β- thfOrflit hanisPh P-rcefla ₈ eofhe ₘ ₁ o5ifa st rfonflu Cleofi ierOfth Ci flatia inlC leoPhi lic„

tt„Ckl ! tuf flror m ₅ dra cYlenzYmtinter mediat ₄ tf ! In thSec ond„ha se>wateri ₛ thfn tpreO fh ile>afldi tt a'-stimefo- Wat ₑ rtfli fr

use to theri ghtsPo t tOrefro mitSnu cle OPhilic„

tt„Ckl ! ti salsonfltaSst ronsanllc leO ₙ hi eas the Sefin ₄ Thepref Orefi, ta kes lo nserpfor watertfrEr form itsfl-Cle OPhili„ atta ₍ kan dbfr eakufac y ! flCn zymf interm Edia t ₜ Undnjitta kes ?fOrSe fiffEtoc ! fat ti tlsfH is57 existsin tOthfhe„

fO„Olh tEdanfd unP fotcolatedfOrmatur in gtheChyflotr Yp sin reac ti O nu ttSp kdOrfO offakes thisPS sibsie n, hePh ySi OlOgic gUPhran g e136l ! nSte adqoraP henYia mni. fl effOiety csilfli„

ar„Ouf-Sua lsusstrfatesofchy motryPsifl) ₁ tSe ani trogfnm cO fhtain infba Si cfOu Pfli„

rt„thelfsll aisubs tratesofttfP sir1 7 ₉ WhatrYp Esof cfmica ₄ reAct ion sarenVo lVeaflmZyfle ₁ 3 7 Theya„tasleflisacidsce rectron- air fCePt ors anlCa flta keparti nenzflec„

ta„YS ismfch-nismfOfen Zymes ls sl F aruelT flEfc nahisflfsofeflzy ₘ at icc at arysisfaretnE sa flCa sphosef nfo Ufut ertaiflfrs anicdemist„ Y?or ₄ fati nsalfa comPlexes 9 cafler a rq ci dCatallSi sisthf rafto ran efl7Ym effcfdni SmfhEre apatli ndc.

dO„Otherm Oleculed On ateSahyd ro genol tpanO thErm Olecule l4fS NiSt„fdsfOrup ₄ noie cuulf Nu Cleo PhilicsUb- itu ₄ iob Theu„im Olecfla ₙ fdr tlfe dnsthti tofesfinstgh-def ki neticsf-fthefCa cti Onisr¥z : +k z +x ₍ wi thdnN rfeaCtiOf-the fa teisae Pe ncel to nth esreedwithwh ichtheExpfakSaw ayf rompterO Th eZ9blrcol esinga.e ranfllic kiyco flna fEdt Othe breflk dOwnofRi bnzStafus rO rtiflOlecu larfluc leOPhilicSubs ti tutionl Th ishappensth ththeS amereactions che meift nezatfac ks thek ₄ -Oie dne beprEithre aksAO wn T hus thfconfe flnr ati flOforlo„ hiExapl: ar eimp ₜ ar ₄ anith-rat eqis ₜ

ys„fcO ndm Ordem ₄ trfes ₄ ur eac ₜ i0 nle adstoms Sofsfereospec ifiCity„ SthEXs ₆ ₜ Pea v esrefor Eth eente ri flg nucle op hie ₄ th isfrean st. ra nthen Uclee Dhi ! Eellenter-flmdiff erentan gie slea dinflUfre fen tisomes rsiza fef„ Su Its dono tP rOv Eth ₄ tth emech anis misCO rr ββ-theCaus Er eSuts fTo meii f Fdentex feri flEn tSc ₀ uldconflrad iCtth eprOfO sEdlle chanism ! ith artda ₍ e ₜ tEmecha nisflOul ₄ have to mOdi lf i edtfla CO mmoda tetlen elfexpef iflle nta reS yusl7 ₁ 7 hatqs th.

Connection between the active Site and Transition State

OO„tfa nsitiors tatea fluo gueOlluhav et oave atefthearalc ar bona tOmvert-h- afliqeCan ₀ OnvlerO Ubwa sOrisinali„

fO„ndb sinfe thtr ansit ionstat tin vo!fe ₄ amolel ta rytEtha Cafa lf-

fll„tw Ouda ₍ olav ₍ tOh aveoly satS OftheSatfCar boflS Ot h ₄ ttt h ere- Olla ₍ flfficiE„ tSfe ci rifCi tfO rmtac ti vESi fE ₁ 4 ₄ ! thfluced fit mOdel assureStfthe en ZYmeflasfthfstrafemufes-Oth ₀ v eafla ₍ hanse ₜ OCO-rOrmtofac lot herrer rfct 1 Yl mqfS ₁ therufi tisn Otbe-

ee„thee flz Yffle ₄ nasurst fatebutbe

tW„en theenZY flfanqtfetrdn ₍ ition state0-the s uis trdte oni tsway to Prod fCt ! dc-ansiti onstuteafa ₍ oSleW il ur i the efizy fle n ifer- inthis flo qeANSwER stf?„ O BLEMSC hap ter1 AfSHlr S ₁ 1 Wa tafl ert easicmem ₍ sfofrh isre Xt? ll -rolYme riSdVer y !d-semOie cult,

her at iOo fL·PO42-HPO4- 1i S2/ 1(PKaofH2 PO4:7|2) , aS-a icU la ted-singsth eH eNd rs0n H.

Se. ba1 Chequa ti n .

2HP. i Sls ouRCe oF t HeBaseF Orm, andH.

1 mUS theadded toco n vertone thirdo fi ttO-heac idform, acc Or ding to t he2/1base/acidrati o weig.

OU. 8, 7 gramso fK2HPO 4(O) 05 moles, ba Sedo na f ormula w. igh tOf17 4grams/MO1 e),dis SO1 vei n sma11q ua1i ty Of dis.

i1. Ed water, add 16 7mLof1M HC1 (gives 1/30f0 o fm OlesO, hYdrO Se ni on, Whi c hcOnve.

tS. /30f the moles OfHP O42-toH2PO4-)a, dd i 1Utetheresul ltingmi Xtur eto One1 iter127 A2/1rati Oo fth eHis eform.

Oacidfo rmi sstill

ne. ded, be Caus ethe pH0f the buffer Stes ame in both P

r. bl eMs NaH2 P04i sa Source0ft he acidfo rm, an aoH ilUs the add edtoco nvert w0 thi rdsofitto thebaseform we i gh0Ut6, 0grams0fNa H2PO 4(0 05m0les, used.

n. fOr mula wei ghto f120gram S/ mO1 e),d is solvei nasm all q uant- ty0 fdisti lled water, add 333 mL0 f1 MNaOH gives 2/30f0.

5. Oles ofh ydr0Xi dei On, Whi ch cOnvert S2/3 oft he0 05 to 1 SofH2PO- toHP. 42-)a ndd i ilUt eTher esu ltingmiXtu retoOne 1 iter 28 Afte r mi Xingt he buf fers0luti.

n. 1 o0m D Contains0 75 M1 aCtiCacidan do 2 5M Sodiu mi a ctate. Th- Pkaof. acti Caci dis3 86 USe the HeNders on. HaSse1b a1che, Uation pH = PKa + 10g[CH3]/[ ] pH :3 86+log([CH3]/[CH3]).

H. 3 86+10g(0 25M / 0 75M) 5 1 12W hat I SthEC On neC tionbetweenT herm0 d Ynamics and L ife? 44 The10 We ring. fentropy ne dedtogiv e ri setobr g a nelles lead stohi ghe rent. OPy in thesur rou n ding s/th USi c reasi ng theentropy of the u ni Ver Se asa Whole 45 com.

a. tmen t a lizati on i n or ga nelles b ri ngs com0One ntsofrea C tionsint Op rOXimit Y with, nea nothen Th e ene rgyc hangeO fther ea ctionisnot a ffected, bu. th e avai la b i lit y0 fCo mponent s a110wsi nto Pr Oce- dmore r ea di1y1 6 DNA Woul dhavehigh.

r. nt fo py w i th t hestr an ds Sepa ra ted Th ereaRetwos i ngl eSt rands- nste ad of0ne d0ubles tran d, and thesin glestrand havemore c.

nf. r Mat i on al mobility 47 se et heanswer to eXe rci Se 43 itisstil l Unli ke 1ythatce

1. Sc oul dhavea fi s e nasbarte yt OP1aS m bUt the questi On 0f p roximit y0 f rea ct ants is m Or e to the point. re t han theene rg Y cha ng e Ofa gi venrea CtiOn 48 It w oU ld be Un li k e ly tha t cells o f the ki nd We k- Owwoul dhave evo 1ve d ona gas g.

a. t Th e lack 0f s0 1i ds an d li quids0n w hich a g f [...] g ate s c ouldf o rm w oU ld make a la rge di ffe re nce 49 T he availab le mate ria1 S. i f f erf rom t ho se that Wou1 dha ve been f ound 0n earth, and con- i ti0n s Oftem pe ra tUre an d Pres s urea re very d iff erent 5 0 Mars, becau. e0f condition s m o re li k e those 0 near th 51 An u mber o fe ne rgetiCa1 1y favOra b1e i ntera c ti Ons d rive the p roce. S0f prot- infold in g ulti mate1y increasing the ent ropy 0f the u ni.

P. 0t 0Synthesisi S end er g0nic, requi ri n g1i gh tenerg Y f rom the su n The c0m p1ete aerobi c ox idati on 0f glu co s e i s exe rgon i c and i S. Source

0f ene rgY fo r many org anisms , i nc1U di ng hum a ns 1 tw 0u 1d be reas- nab1e to eXp ect t he tWo proces se s to take pla ce di f ferent.

Yi. 0r. e rt0 prOvid e ene rgs yf0 r the e-dergonic on e Chapter 2 Ans wers 2 2w hat i s a Hy dro gen B0nd ? 1 P r0te inS an d nuc le ic A cidS h ave hydrog.

nb. nds a s an impo rtant part of the ir stRuctures 2, Repl iCat io n 0fDNA an- i t St r ansc ript i0 n t o RNA re q ui re s h ydrogen bondi n g of c0mp1em ent- r Y base s to th e DNA t empla te s t r and 3 The c. Hb ondi snotsuff i cient 1 Y po1ar f or s t rea t1 y une qua1 d istri bu tion f e1ectro n s ati t stwo e nd S A1SO, t he r e are n0 u n Sha red P ai rS o f ElectronS- 0se rVea S hyd ro genbond aCcep.

0r. 4 Many m0 1eCU 1es can f0r m hYdro gen bond S Examp1 es li g ht be H2O, CH3 OH NH3 5 F0r a bo nd. 0be ca1le d a h yd r0gen b0nd, i t mus t have a hydr0 gen cov a1e nt1y b0 n ded t o O, N , 0r F Thi s hydro gen the n fo r ms a hyd roge n bo nd wi th a not her 0, N, 0r F 6 Inahyd.

0g. nbonded dime r o f ac eti c ac i d the -0H p o rti on of th e ca rbo xy1 gr- up 0n m01 ecu1 e 1 i S hydro ge n b onded to th e -(=0) p0r ti on 0f the. ar box Y 1 grou p 0n m o1ec U1e 2 , and vic ev ers a 7 G1uCos e 1 7 and so rbi t01 15;e a C ha 1Co hO1 gr0 u p Ca n bon d t o water Sa nd the r i ng oxy gen bi n ds t0 t wo T hes Ug.

ra. Co ho1s bind mOr e tha n t he corr esp0ndin g suga rs) 8 HY po vent- 1ati0n a ccre aseSthe p H0 f b10 0d (see BC onp 54) 9 A SPi r ini- e1eCtrica 11 y ne utral a t the pH of the st0. ac h a nd can pass t h ent m b ran- mor e eaSi1y t he r e th an i n the sm a1li ntes tine 1 0 P 0Si tiVeIy cha rged i ons wi11 bind to n ucle i. aCi d sasar esultofe 1ectr 0s tatiC attract ion t other egat.

Ve. Yo a rge d Pho sphate 11 T h e un i que ri t n es S of water f0 r min g h ydro gen b0nds de. ter mines the propert ie sOf many i m portant bi 0mo1 eC u1e S wate r c a n a1S0-c t a s an aci d an d a ba. e, g [...] gi tgr eat Ve-Sa ti1 ity in b i0chemica 1 rea cti0ns 1 2 Ii fatom S did notdi ffer in e1ectro ne gativi t y, the r e wou1d be n0 pol ar b0nds T.

iS. 0U1d drastic a1 1y a ffect a 1l r eacti 0 nS th at invo1ve fu n. ti o n a 1 grou ps c0nta i ni n g 0xyge n or ni t rogen That i s m0s. 0f 2 3 Wha t Ar e Aci d s a n d Bases? 1 3 (CH3)3 NH+ (co n jugat.

ac. d (C H3)3N (conjugate ba Se) + H3N-CH2-CO O H conjuga te a ci d) + H3N-CH2 -COO (conju gate b ase) + H3N-CH2-COO(c onju gate aci d) H2N-CH2 -COO (conju ga te b ase) -CH2-COOH c onjugatea Ci d) -CH2-COO(c onj ugate ba Se) -CH2-COOH(conjugate base) -CH2- Co OH(conj ug.

te. C i d) 4.15 (a) The numerical co nstant eq ua l to t he c on c entrati On of the Pro ducts 0f th e di Ssoc i atio n d iv i ded by th. con centra ti0n of the u nd iss0ciate d a Cid f0rm: ([H+][A-]/[HA]) (b) The equalit.

ti. e0r qua nti ta ti ve descr iption of how mu C h a Cid (HA) d i S s o c iate s t 0 hy dro geni. n (c) The pr0per t y of am0 1ec U le t ha t has both a po lar regio n and a n0n P o1a r r eg ion (d) T h e amou n t of ac. d or ba se tha t can be a dded t0 a buf fer be fore e Xperie nc i ng a sharp pH c hange1 (e) T- e p0i nt i n a ti t ration c urv. W h ere th e a d d ed a ci d or ba se

qu. 1s the a mount of b uf fer origina11y p resent (f) The p. Ope rty fam ilie d th at i s f ea di 1yso lu b1e in wate r(i le 4 5 [...] whic h in t u rn ha S1 ot i mes 1 es S bUff. ring ca Paci tY t ha n the thi r d A11 thre e buffers h a ve the sam e pH s ince they a1l ha ve th e sam e re1a ti ve amoun ts of the ac id an d ba Se f0.

m .5 It woul d be n0re effe ctiv e to st ar t w ith the HEP E S base y ou and b

hece lu L YSosomes con t ai n hydrolytic e n ZYm es, per 0xi s0m es cont ain ca t a1 aSe needed for th e me tab0li sm 0f per0x ides), and gly0xy S0 mes con tain en Zymes needed by pla nts for the gl Yoxy.

at. CY c le A 1lth ese 0 rgan e1 1es have th e ap pea ran ce of flatt ene d sa cs boun ded by a single membrane. 1 7 H 0w do we Cl aSs ify org an is ms: Five k ing d o m s or Three 2 s M one ra i n c1ude ba Cte r ia e g 1) E coli b a nd cyanobac te ria Proti st a in c lude Eugl en.

V. 1Vo x, Amoe-a, a nd Par amecium F Ungi i nclude mo 1ds and mushroo m s Plan ta e inc 1ude C1ub m osses a nd oak trees An i m a1 i a in c 1u d es Pi d ers, sal m on, rat tles na kes, r obi ns, and d og s 2 The ki ngdo m Monera C0n Si. ts o f Prokary 0t es The 0the r f0ur ki ng d 0ms cons i st 0f eu ka ryote. 3 0 T h e ki ng d om M one ra i s div id ed i nto the do mai nS E ub acteria and Ar chaea on the basis o f bi0ch em iCa1 di f f e. enc es The do mai n Eukarya c0 ns iSt s 0f the f our k in gdo ms of euk a ryo- i c orga ni s ms 1 s Is there co mmo n gro un d for a11 c. 1 is? 3 Them a j0r advanta ge i s that o f ha v i.

gc. mPar t me nts(orga ne1 1 es) with spe Ciali zed func t i ons can d th- s di visi 0n 0f la bor. A1s0 the ad van ta ge i s that cel1 s can b e much l

rg. rWi th su ch a re a to v0lume co nsideratio nsbe i n g critiCal b ec a-use of co mpartm ent a 1 3 2 S ee the discu SSio n of the endos Ymbiotic.

he. r Y i n Secti0n 1 q 33 see E XerCise 32 T

he. iv i si0n o f labor i nce1 1s gives ri s e to greate r effici enc Y a nd a l arger nu mb e r of ind i Vi dua1S Th is in tu rn a11 0ws m ore 0 p portunit y for e v0 lu tiona ry s peci a tion 1 How Do Ce 11s use Energy 34 P roce- sses that release en ergy are fav0 re d 1 1 0 What is t he co nnection bet.

ee. Ener gy and Ch a n ge? 35 The term s pontane0u S me ans ener ge tica- lly fav. re d It does not ne cessari ly me an fa st 1 1 What i s t he crite rion for spon ta. ei ty in biochemi c al 36 T he s y stem i s t h e non- polar s0 1ute and water, whic h become more di sor dered w hen a1so.

Ut. 0ni s for me d; s S ys i s P0 siti ve b u t c0m parative 1y sma 1l, the ss urr o undin gs i s nega ti ve a nd co mpara ti v e1 y la rge b e cause it i s a refle c ti o-

ft. eun fav Or ab1e enth a1 py Change for for mi n g th e so1 uti on (H 3 7 P roce sses (a) a nd (b) a re spo ntane ous.

Wh. rea s Pr o Cesse S (c) a nd (d) a re n0t Th e s ponta neou s Pr ocess es rep res ent an in cre ase i n disor der (inc rease . n the entro p y o- the Univ er se) a nd ha ve a neg a tive G at c0nstant tempe ratu.

ea. d Pres Sur e, w hi le the oppo Si te is tr ue for th e nons pontane0u s pr0 ce S ses 38 In a p1ca

Se. , the ri se in Crease i n ent r0 py , a nd t he f i na l sta t e h a S more poss i- ble r. ndom ar rangements tha n the in i tial sta t e 39 Si nce the equa ti on i n vo1 ves mul ti p1iC ati0n 0 f S b y T, the va 1ue 0f G i s temp. ratu re dep endent 40 I f one c o ns iders e ntr O p y to be a measu re of dis pe.

Si. n o f energy th en a t high e r te mperatur e s i t i s 1 0 gi Cal t h a t m0 1ecu1es w 0 u1d ha ve m or e po ss ib 1e a r ra n ge men ts d eto i ncr ea sed mo1ecular m 4 1 Assuming the v.

1U.0 f b s i s p0s i ti V e, and i nc re aS e in te mperature Wi 1 l i n C

ea. e th e co nt ri b ution of th e entrop y c ompon ent to th e o ver a1 1 en e r gy ch. nge 42 T he h eat exchan ge, e t th i ng c0 1d e r, r ef 1 ect s 0n1y the entha lpy c0mp.

ne. t0f the to tal energy cha n g e T he entropy change mu st be hi ghe r to ugh t0o f f s et the enthal py comp on en t and add up t0 a n over a l 1 G 43 Entro py would i n c r ease The rear.

m0. e W ay s that two m0 lecu les , ADP and Pi , ca n be r a nd0miz e d th a n a s i ng 1e mo1ecule, ATP 13 [...] Wate r mov ing (S) the p r0p-

ty. fa m0 1ec u1e that is i n s0 1ub1e in water c (e) water (b) T he pr0per ty of a mo1ecule tha t i s n0 t s0 1ubl e i n Wa ter) The p ro p erty o f a c ova 1e nt b0 nd where th e r e i s eve n sha r. ng o f e 1e ctro ns an d n0 di po1e m0me nts (par t i a1 c h a rg e S) (i) T he p r o p er ty of a mo1e cul e that i s S0 1ubl e i n wate r the pro Per ty 0 f a cova1e nt b on d whe re t he e lect r0 ns a re n0t sha red eve n1y a n d dipo1e mo me nts (par ti a l cha rg es) exi st (j).

ne. Peri men t whe re bas e is added to a n aci d or base, i s ad ded s0 t he p H wi s et a c0m p0und and the pH i s mea s ure d as a f unc ti 0n o f th e adde d s ub 2 14 Wh at i s pH- and wha t d 0es i t hav e t0 d0 w ith the Pro pe rtie S of wa ter? 16 T he de fini ti 0n 0f pH.

S. 0g [H+] d ue to the 10 g fu nctio n, a ch an ge in c0n ce ntrat i on 0f 10 Wi 1 1 1e ad to ac ha ng e i n pH 0f 1 Th e 10g 0f 1 0 iS 0ne Th e 10 g of 100 i s 2, e tc 17 B 10od p1a Sma, pH 7 4 [H+] 4 0 X 10 8 M 0ran ge jui ce, p H 3 5 [H+] 3 2 x 1 0 4 M Hum a n ur in e, p H 6 2 [H+].

6 3x10 7 M Ho use h0 ld a mm0nia , pH 11 5 [H+] = 3 2 x 10 1 2 M Ga s tri c j- ice, pH 1 8 [H+] = 1 6 x 1. 2 M Sa l iva, pH 6 5 [H+] = 3 2 x 10 7 M Int race1- u1ar f1Ui d (1i ver) , pH 6 9 [H+] = 1 6. 10 7 M To ma t0 ju i ce , pH 4 3 [H+] = 5 0 x1 0 5 M G rapefru. t ju ic e, pH 3 2 [H+] = 6 X 10 4 M 19. S

1i. a, p H 6 5 [OH] = 3 2 x 1 0 8 M In trac e1l u1ar flUid (1i Ve r), p H 6 9 [OH] = 7 9 X 1 0 8 M To mat0 ju i ce, pH 4 3 [OH] = 2 0 x 1 0 1. M Gra pef ruit ju iCe , pH 3 2 [OH] = 1 6 x 10 11 M 2 16 Wh at Are Buf fers, a nd w hy Are They Imp. rtan t? 20 The pKa of th e buffer shoul d be c lo se t0 th e desi re d buffer pH, a nd the s ubs tance shoul d n0 t i nterfere.

it. the rea Ction bei ng 21 The use fu l pH ra nge of a buffer is. ne pH uni t a b0ve a nd be l0 w its pKa 2 2 Use the Hend erson Ha sse1ba lc heq

at. 0n p H = pKa + 10g([CH3]/[CH3]) -0.4 7 6 + 10g([CH3]/[CH3]) 4 10- ([CH3]/[CH3]) = i nve rs e -0g0f0 24 1 7CH3C0 0H 1 2 3 Us et he Hende rson Ha sse1ba1c. equati0 n pH = pKa + 10g ([CH3]/[CH3]) 4 00 = 4 76-

([CH3]/[CH3]) = i nve rse 10 g0 f 0 7 6 = 0 1 7 [CH3] 24 Us e th e Hende rson. a Ss e1ba1c h equ ation p H = pKa + 10g ([TRIS].

+ 8 7 = 8 3 + 10g([T RIS]) + 4 = 1 0g[TRIS] [TRIS] i nvers e10 g0 fo 14 = 2 5 + 1 25 U se the Hen d ersonH a sse1ba1c h equat- on pH = pKa + 10g([HEPES]) + 7 9 = 7 5 + 10g([HEPES]) + 0 35 = 10 g[HEPES] [HEPES] = i nverse 10 g 0f 0 35 = 2 2 + 1 2 6 at pH 7.

g(.J Oo2 /o I O0 8)PH= 7 7035 Fif Stcalcll

1a. et hemplesd fbuffer, tha ty ou have: (w eq regoing to dOSometro un din gofr)1 ooml=.

11. it er s ando, 1 lit er, of 0 1 M TRI Sbuf fer is o 0 1 mol es (Si nceth e bffe fi satpH7 7o, w e saW in exercise 25 that the amount of TRIS iso o2mo1 es) and t hea mount of TRIsH+ is 0 0 8mo. Es If you then ad d3mLof1 M HCl you w ill b ead ding o 003mo1 e s Of H+ T hi s Will eact as Sh own: TRIS+H+→ TRIsH+ U ntily Our Uno-tOfsom ething Whi ch W i11 et he TRIS S ince it is the l imi ting rea gent al 1 the TRIS is c onverted to TRIS H+: n=o 0 m

eS. RIS: No mo1 es we hav e us ed up the bu ffer C apacity of th e TRIS we no wha e0 0 3m o1 eS of H+ in appr ox im ate1y 0 11 tetS of S01u ti o n T-is is appr. Xi ma te1y 0 3M H+ pH = 0 go 3pH=o15 236 [H+]=[.

]. Or pur eacid t hus K a= [H+] = [ax] HA 21 0s [H+ 1 PK a 0s [HA-pH=. /2{ PKa log[H A] 3 7 use the He nd erson Hassel balc h equ atio n [A] te ate iO n 1 /lace. ic a Cid 1 2 3/1 38 Subs tan ce wi th a pKa of 3 19 as a buffe r ang to f21 9to4 9 l t w i 1 lnot buffer effe C tiv e1 yat pH7 5 391 use the Hender son H asse1.

a1. he qu ation The ra ti o of A w Ould be 3 8 1to 1 401 in al 1 Ca- es the slightable. Uffer ra nge Ove r a p H r ange o f pKa+/-1 PH uni ts. Lac ticacid (P Ka=3 86) and its sodi u ms a1t pH2 86 4 8 acetic acid (pK a= 4 76) an d itsSodiumSalt pH 3 7 6 76 TRIS (see Tabl e3 4) pKa= 8 13) initsP rOtO nated fo rm an d it s free amine form pH7 3 9 13 HEPEs s. e Table3 4) pKa=7 5 5 init s z w itterion ican d itsan ion ic for m pH6 5 5 8 5 54 1 se ve ral of t he buffers would be sui tabl e) ma nely TES HEPES MOPS a nd PIPES.

OWever t he best buf

fe. WOUld be MOPS as i ts pKa of 7 2 i s cl oses t to the desi r

e. PH of 7 3 -2 The s 0 lution is Sca 1 led to 500 molar eVen thou gh the co nc ent rati on 0f one i ther t h e free base no r the c onj ugate aci d is 0 05 M 0 5 M the correct Concentra tio n to report B. ffer co n C entrati ons are.

yp. Ca 1 1 y reported to be the sum of t he tw O forms (1 43 A t the equ

V. 1 e qui po i nt Of the ti tra tion as ma 1 1 a mount of a c et i c aci d r em ain- because of t he equ i l ibr ium CH3COO H ⇌ H+ +CH3COO The r e i s a sma 11 a mount of acet ic acid le ft 44 bu ffering ca pacity is a se d upon the a mounts of the acid and base forms present in the buffe r solution A solution with a high

b uff ering capacity and pH With 1 arge amount of acid and base with a

S t i c Change s in pH A solution with a 10 W buffer in g cap a

C. t c an re act W ith only co mpara ti ve ly sma 1 1 amo u

t. O f a Ci o r base before sho w i ng c hanges in pH.

Th e mo re concentrat ed t he buffer the higher its buff ering capa city The first buffe r is the.

h. re h a s lo ti mes le ss buffe ri ng Ca pa ci ty than the s e Cond giving the c arbon d part. a double bond c haracter 30 1 I 11 - ase s t he yield s 0 95 n For o residu e s t hat m -ar 60% yie1d f-

r50r. Si d ue-) g and f or 1 0 0 res i d es 0 6 % These are t o tat is f a -toyi elds En zyme spe ci fic i ty get ar ound th. Pro blem 3 1 The to pe pt i des diff e ring in n o C ids equence bu t n ot in co mp o 32 Th tit ration cur.

es. f the two pe ptides will have these the same shape

I. he pKa va1 ues o fth e amin o a nd car box y 1 groups will differ ver y carefu l work will show S li ght differences. Each amino acid val- es be caus e of th e differe nt di stan ces to the ch arge d groups at th e en.

S. fthe Pept ides uch anges are par ticu lar ly marked in pro tein

I 33 AS Pl Leu Phe le UaSp Phe Ph e ASp Le u ASp Ph e L eu Leu Ph.

Sp 34 D LFL DFF DLDFLLFDFL D 5 yellow ou1 d get 2 0 2 x 10 m o1 ecu les whi c is about the earth vo1 ume s t he same. Calcu la tion of a p enta pe ptide s more comprehensible res ults 36 The ya re rat ive ly stable they a.

ezi tte rio ns the y typ ically have h

i. hile i t in g i ts 37 with α-rylit the doubt no I c ompare P re dicti-g the pro perties of water fr om thos e of hydrogen and ox. g n init he rat o i co r m o le c l ar form If y0 u knew the p.

O

P

er. i es of the p rotein you might be a ble to do the rev erse to so me e xten

3 8 These two pe pti d es differ c hemi cal 1 y. T h e open chain has a free c arbo xyl te rmina1 and N termina1 b ut the cy cli c pepti de has on1 y

Peptide 3 9 0 t he c te r mi n a 1.

nd. he N termi na S.

ft. eOp en chai n pepti de c an be charged at app rop r ate pH values Whi chi.

no. the Case wi th the.

yC. icpeptide Thi s can provide a basis for sep arat i on by 40 carb-hydrates are no t as our Ce of t he nit ro gen neede d for biosynth es is of amino acids 41 sugg est that you.

fr. e has both e α-ca rbo xyl gr α-U p as a charged carboxylate

(. Oo) an d the amino group in its c harged form N H3+) 4 2 he. ear ever fews i de ch a i nsth at have funct.

On. 1 group stor or m 43 There will be a n y more possib.

ec. n formation s be caus e Of f ee ro. at ion aroun d t he pe ptide b ond 44 Ther e wou 1 d be no Possib ili ty fo is ul f-decrossli n-

rata p H a b ove the p Ka which means that th.

ba. e from Wi 1 1 predomin ate When you have

fi. iS h ed prepari ng it It is e asi er to c o ne r tson e of the bas e form to he a ci d fo rm than most of the ac.

df. rm to th e base for m 46 I n a buff er with th e pH above the p Ka, t.

eb. se form predomi nates this would

be. S e fu -as a bu ffer for a reaction that pro d uces H+ be cause there will be plenty of the base form t o react with the hydr oge n io n produ. ed 47 Z w itte rion s tend not to i nterf ere with bioch em i cal reac.

i0. S' 48 It s use f u l to ha ve a b uffer that will maint a i n a Sta ble p H even if a ssay C ondi ti ons ch ange dilu ti ons uch po ssi ble chan ge 49 It is useful to h av e a buff er that wi 1 l maintai n as tab le p H e ni.

aS. ay c ondi ti on s ch a nge T empera t ure va ria tio n is on e su ch poss ible chan ge 50 Thi s 1 Zw itteri on i S+H3 N-CH2-COO C hapter 3 A nswer s 51 what a re Am i n o acids and Wh a.

iS. he i r three -di mensional di a nd L am in o ac ids ha ve di ffer ent ste reoChemi. try around the a c arbon pepti des th at con tain D-amino acids a re found in n a tural ce 11 wa 1 l sa n d i n some anti bio t i C s 3 2 What are the s tructure s a n d Prop erties Of the m.

iV. dual Amino Acids 2 P rotein ei ste C hni c a1 1 y n o t an amin o acid 1 y cine con tai ns no chiral c arb on at o ms 3 1 A nami

Oa. id in which the R g ro u p contai ns the fol lowi ng a h y -roxy 1 groups eri n e thr eo nine t yrosine) a S u1 fur atom cyst ei n.

m. thio nin b ase Con d ch iral C ar b on a tom c is ol eu c ine thr eo n ine a n amide gro up (as p a ra gi ne glutamine an a cid grou (as p.

rt. the glutam ate an aromati c ring phenyl a la ni ne ty rosine tryp t oph an b ra nched side chain (1 eu cine vali ne) 4 In th e pe ptid e val.

Me. er ple he lar ge ys ary r e u th e p olar amin o a cid s ar es e A rg ys and T yr the aromati c amin o acid s ar e phe and Tyr an d t he sul fur C on taining am in o acids. re Met and cys. 5 1 n the pept ide glu Th r Va l S e r Ala te n o.

PO. ar a mino acids are Va l Ile and A sp t he acid ic amino acid s are Glu and Asp amino aci ds other th.

nt. ell s are produc ed by modi fi cati on of one of the common a mi no acid s se. Figure 3 15 for th e s truc t.

re. Of some m odified am ino a cids hydroxyp rolin e and h y d roxy l ysine are fou nd in col lagen th yroxin e is fou nd in thyro globu lin 7 Ty r osi ne try ptophan and t hei r deri va tive s 8 A mon a mine oxi dase is an enzym e that deg rades c ompound s w it h a n am in o.

ro. P i ncludin g neurotran smitters co nse quen tly th ey can c ont-ol a per s on s mental state 9 Th e high con cen tration of tryptoph an i n milk protein ma y m ild ly e l.

va. e the l-ve l s of s erot onin which relaxe s the brain 10 The. rypt oph a n in m i lk mi ght make you sleepy w he reas t h e tyram i ne in cheeses h ould pep you up 11 The a min o aci d th yroxine and-

ydroxyproline occur in very few proteins The s.

ne. iC C o deO es not in c lu de them so th ey are prod uced by modifica ti 0 n of tyr os i ne and pr oline respective ly 31.

DO.

in OA cid s. Ves pecific aci d-base Pr oper ties 12 14 Histidi ne: i midazole is dep r ot onate d amino group is pr edo minantly deprot on ated As paragine: amino group is deprotonated Try-topha n: a mi no group is p re dom in an ti y depr oto na ted pr o line: amino group is p arti al ly dep roto n at ed ty ros i ne: amino g rou p is pre dom inantly deprot on ate d the phe nol ic hydr oxy l is approximately 50 % o f fix. Ure o f p roton a te d a nd deprot onated 15 I soe1ectri c point for glu ta mic aci d 3 25 s e rine 5 7 is ti di ne 7 5 g ly S i ne 9 75 ty ros ine 5 65 arg inine 10 75 I 16 C ySte ine wi ll have no.

et h ar ge at pH 5 1 o2=(1 71+8 33)/

2(. ee ti t ratio n cu r ve he b) 7 pH= 3 86+(lo 48) pH=3 38 29 After mix ing the b uffer s o lu tion c 100 mL C on tains 0 25 M la C ti c a ci d and 0 7 5 M s odiu m la C tat e the p Ka of la C ti c aci d is 3 86 Use the He nd e rs o n-Ha ss e lba l Ch e qu at ion pH= pKa + lo g [C H3]/[D 3 pH=3 86+lo g [CH3][D 3 pH=3 8.

+1. g(0 75M /0 25 M) pH=3 86+(0 48) pH= 4 34 30 Use the H ende rson-Has se l ba l Che q ua tion pH= pKa+ lo g [CH3][D 3 pH= 4 7 6+10 g[CH3] D CH3 pH=4 76+ 10g(0 25 M) (0 1 M) pH=4 76+ 0 40 pH= 5 16 31 Ye s it is co rrec t calcu l ate t he m ola r amou nts of th two form s a nd i s ert in t o the Hen derso n-Has se1 b a1c.

eq. ation 3 o2 g=0 01 6 7 mola nd 5 25 g=0 033 mol) 3 2 The so lutio is a buffer b e ca use it co nt ai ns e qua l c.

nc. ntra ti o ns of TRIS i n t h e aci d an d f ree amine f orm. When the two solu ti ons are mixed t he c oncentrat ions of the resulti.

gS. lut ion c in the a b sence of re action a re 0 05M HCl and 0 1 M TRIS because of dil ution T he HCl re acts with ha 1 f th e TRIS pre sent giving 0 05 M, R IS protonat ed form and 0 05 M TRIS free amine for m 33 Any buff er that has e qual con ce n tra tions of the acid a.

db. Si c fo rm s wi ll have a pH equa l to its pKa th er efore the buf fer f ro exercise 32 will have a pH of 8 3 34 First c. 1 Cu late the m ole s of buf fer that you have: 1 0 0 mL 0 1 liters and 0 1 liters of 0 1 M TRIS buf-fer is 0 1 mo. es Since the buff er is at its pKa, the r e are e qual co nce ntra ti ons of the acid and b a si c for ms so the am ount of TRIS is 0 005 mo les and the amount of TRIS H+ is 0 005 mol es If yo u the n add 3 m L of 1 M HCl you will be adding 0 003 mol es of H+ This

Wi. l rea Ct as Shown: TRIS+ H+ → TRIS H+ Un

ti. You run out of somet hing Whi ch wi ll be the H+ since it is the li mit ing reagent The ne w amo unts can be cal cul ated as sho wn below: H+ = 0 005 mo1es + 0 003.

01. S= 0 0 8 mo 1 es TRIS 0 005 mo1es 0 003 mo 1 s= 0 002 mol es Now pl ug th e se val ues in to the He nders on-Hass.

1b. 1 Che qua tio n: pH= 8 3 + 1og([TRIS]/[TRIS H+]) = 8 3

01. gy i smarg i nalforUSe Ofcomparati, eMo de li ng Iti Sbe st to tr ythatmetho d, bUtt he ntOcom, areth ere sUltsw it hthoS eOta iNe dfro Ut reroldrecog ni tion 36lse ethePrOtei NatagaNk 37.

AP. ionisap Otent ia11 yi nfeCtio usProtei, founainmu it iPi eforlsi. Mamma ls, Often con cent ratedinN rvoustissle T hefei sanor al forman danabnorm dif ornt hattelast. forlplaque sdestrOin gthene, lo uStissue T hey have been fOu ndtoretralls mi Ssib lelcrss, peCie Si 3 8 Aseries ofence Hi loa thieshav ebe enfolll.

t0.ecaus edb Ytheabn0rmalPri OnProteinl1

C0. S, th ediseaSecau sed byPrionsiSc a1ledbovi nes pongi for Ilencepha lo Pat hyormo rfco mil onlymadCowdiSea se I nshePthediseaSeis. a1ledsCrafi lt In humansi tisca 11edJak ob Creltzfe 1dd iseasel 39, The nOrma1forb ftheP rion PrOte inhasa hi gh erhe1ixc0nt entcomp.

re.t0 bsh ee tl TheabnormaI on ehasa

ni. Cleased 47 whaitheQu ate mar'St fu Ctu reofProte. ns ?40 b OthCOnta inlem, groll p; bothar eoNg. nbi nding Se COn darystrlct ur eiSPrillarily, heli, D iffe rence.

Sa t e tr allen jlemlog 1obin, sanO no ller;o xy gen bin inglOHemog lo biniscO Op erative, butnonco operat.

Vetomyo gl Obin 41

T. eCru Cia 1re Sidu esafe histi dj n esi nbothPr

0. eins 42 S hig hes tle velofolga ni za tion iSte rti a ry Hemo. 1. bin Si-qua, ernary 4, The -nctionof hemog, obinisOX-gen, ransPortits, igm oidaibi ndi ng curverefle, tsth ef actthat i- canbi ndea si 1Ytoox yge n-tCompa rnt.

V -Yhis hpressu resandre1 easeoxy genati -er Pre, ls ure S Th efunctio lb fmyoglo, in isoxyg en storage; asaresl1t, i ti-e-si 1ysatur-te dwi thox ygen atlo- Pressures, as shownbyit-hy per bo1i chindin curve 44 IntheP- esenceofH +an dCo2, bothofwhi chbi1lat 0-emog10bin, theox- en bildin gcapac i tyofhemoglobindecr- ases 45 Intheabs en ceof2- 3 bisphosphog 1ycerate - thebindingof, Xy genbyhemog1obin, eSem b1 est hato fflyoglob in, Character- ze dbyaCkofcOoperativ it y 2,3- BisPhosphog1y ce rat ebin dsatthece, te rofthehemo globi nmo 1ecu1e, i -re ase Scoo Perat ivity, St abilizes thed eoxyconforma, iono fhemo g1ob in, andmodu1a, es thebinding ofoxy gen sOthatit α- -ea si1 Yber eleasedint hecapi1-rie- 46 Feta 1hemo glob i nbin dSoxy ge nmores trong, y tha nadu1 them Oglobin see Figure 4 -5 47 Histidin e1 4 inal Chain is repla- e dbyaserine ina 4, deoxygen atedhe mog1obi nisawe a ke racid (Ha Sahig herP Ka) t hanoxyg enated hem.

g. Obin In Othe rwords, deo xygenate dh emoglo binbinds mor estro- g1ytoH+tha, dOe Soxygena tedhe mog1ob in Theb in dingofH +and ofc- 2 to hem, globi nfav orsth echang ei nqu ater, ar y s tructure othe deoXygena ted formOfhem oglob.

n. 4 TheP rim ar yfla winyourfriend s r eas oningi sare versa1 Ofthedefin itionofpH, whi ChiS pH= 1og[H+] I fthere 1ease Or bind-

gfh. dro geni0nbyh emog10 bi nwere t-ePr im rifactor i nthe Bohrefrec tthe PHCh ang eswould hetheoPPo, iteofthoseactua 11y Obs erved The res p o nseofhemog1obi nto changesinP H, sth. cen tral POi nt- henthePHin cre ases, thehyd rOgenionconcentrat- 0 fdec reases, andV i ceve rs a 50 TheC, alsgeora- i-ti dinetoa serin eint he. chai nfeN- les a9os i ti Ve1 ycharge aminoaci, tha tcOUld hav eintera cted with B pd Thus, t he rearere lers altbr. dgeStob reak, sobindingisea sie rth anitisi na 51 ler sOSwithsic kle celit raitha ve sOm eab- Orma 1hemoglob i n Athi gha1tit lueY ther, iS1es sOxyg en and- e oncentrat ionofthed eox yfOrm oftheabn. rmalh emog1Ob i n increase, lessoxy gen an, ulati cau singthe, erV edb e a t hingdiffi cult ies 52 Infet alhe moglobi n, tleSubuni tc0 mbs ti0nis 22 Wi threplacement of thechainsbytheChai- S Thesi Ckle cellmutati onaff C, S- he cbainsot hef etus- omzy gou-f. rHbshasnormalfe talhe mOg1Ob in 53 The re 1 ati veOx- gen affinit ies a1lowOX, ge ntobe tak enbythe fetalce 11sfrOmth emate rnalH 5 4 BecauseP eOple with SiCkle ce1 ldi Seasea.

e. hron ica 1 1yane, ic, S om ecelIswithfe, a1Hb a reP, oall ceat0h- 1rO, eFco m ethe im pa ir edo xy gendlive. Ysys te m 5 5 The, rys tal 1ineform ch ange dbeCause oxy gen enterea nderth e0 overs1. P, tr ansfo rm in sde oxyhem og10 bintoOX yhemOglObin cha.

t. r5A nsWers 51 1 HowDowtext ractPure pro te insfromC. 1ls? 1 B1 inder, P otterlEiNe ine m'sonicator 2 lf, ollneeded tomaintai nth estructur, i n te g rit Yoft hesubCell ularorg. ne1 les, alo tte r e ive jhemOu 1dbebetter. ec ausei tisMOre gent le The tiSSue, SIC has1 iver, mustbeS ofte noughto usewitht, isde Vi cel 3 saltin gouti sa.

r. Cesswhere byahighly ion icsaltiS ed toreduc eth esO1ubi lity ofaP ro teinuntil itc, mes0 utofs0 lutio nan dcanbecentrif uged The a1t form Si On d ipo 1ebondswi. hthewat e rinthesolu tio n, Whi ch leaves 1es swaterava i1 abl etOhydr ate t he Protein N onpolarSi. echains be gi nt ointer ac the tWe enProtei nmol ec ules, andthe ybec Omei nso, ub1 4 Thei ra minoacidC Onte nt andar rangemen tsma ke sOm eProt eins mO res o1ubleth.

n. Ther s APro te in w i thmOre hi ghlyP, la ra minoaCid Son thesur facewi11b enOr eSo1u bl etha n one wi thmoreh ydro. hO bico nSon thes urface 5 HOmogeni ze th eliverce1 is usin gabott er 1 ve jhem HOm ogen i ze r sPi n the HOmog.

m. eat 5o0oxgtoSedime ntt heUnbroke. Ce11 s and nu c1e i Cen, ri fu get- esuper nata nt at1 5, 000xg, an dco1 1ect the P e11et Whichco.

ta. nS th emi tochondria 6 NO, Pe roXisomesa ndmitochon

dr. aha ve0 verla PP insedi ment altionCharacteri sti Cs o ther tec hni q ues Suc hasSuCrOsegra dien tcen tri fug ati. n, w ou1 dhave t0 beusedtosePa ra tet het woorganeIles 1 invO1ve din c0 king mea tar. Usual1y morethan ne nous hto denat Ure th ePro te inP art ofthe Me, 9 Pri Ondiseas e shav ebe en1ink ed tOth. immuneSys tem Iti s e1 iev edthatth eP rionProte inst ravel. nthe, Ymhs ystem bolulat o1ymP hocYtesand e ven tua1 1yarriV.

at. hene rvO ustisSuewhere the Yb egintO tran sfor mth enorma 1P ri onProt. i nin tot heabnO rm a1One 10 Whi1eth ere areStronggene tic PrediS po sitionst. acqui re, crapie t hata1One Wi 11notc aus eth edisea. e T hedis easemis the ta rtedby inge. tin gaPri

etween Pept i d echains, gi ving morePo ssib1e cOnformati Ons I T herewO ul dnot be t herP os si bil.

ty. foXidati O n red uctio nrea C ti onsinVolvi ng U lfhydry1a ndiSu 1f ide, r0UPS 45 Theb igdi f fe re nc ewOul dbet he 1os S ofst ere Ospec ificity.

nt. econfo rm a tionofa nype ptideor proteinTh. SW ouldhav edr astic co nsequenc esforthe ki n ds offeacti onsOfthePr Oteinl 3 5 ArtsmallPep tides PhyS io1Ogica11yA Ctive 46 O Xytoc in a sanisO1ecu.

in. atP, SitionSandl euCi ne atP Osit, on8, its ti mu1 atessmoot hmu Scle con-tractio ninthe.

te. UsdUri ngiabor and int hela mmaryg landsd. ringla Cta tion va- 0Pre ssi nhaS aP en Yla1anine at Positio n3and arg in ineatP- sitions, i. Stimu latesreso rPti On ofwater byt hek i dneyS, th uSr a

iS. ngb lood 47 Th ere a ure dfrom Ofg 1lutathi onecOnsist sof thr eea- i noaci dS Witha Sulfh, ylgroll, ; th eoxidizedf0 rmC OnsistsOfs ixaminoac. dSan dCa nbeco nsider edt here Suitofl i nki ngtwo molec ules0, re duc ed glu tath io nebya disulf ideb r idge 48 En- ephali ns arepent, peptides(Y-G-G-F-L), leu cine nke- halin, and M.

F. Methi O ni n eenke- halin, whichar, natur a11 yoccurring an a1g esic 49 Inmos t Cases, Damin oac i d Sar et.

Xi. Th eyocCu ri nnatur ei nan tibi otics andbac teria1ce11Wa11 s 5 0 Thediffe renstse rdoch emistry0. thet wopePtide S1e adstodi ffer ent- indingWi thtaStetece PtorSand totheS we ettastefor orean dthe.

it. ertaster Ortheo ther Ch apter4A ns Wer s4 1 HO wDOes theSt, uCtur eo fPro teinspe ter mine Their FuN cti On?2 (a)<(2); (b)(4); (c)(b);(d)(3) 1 when aProtei ni s denatured, theinte ra cti onstha tdet ermin ese condary, tertiary, an dquaternar narystr uctureSare, Vercolleb YtheP r esen ceo f thedenaturin ga gentu, 1 Ythe Prim arYstru cturerema i nsi ntact 3 Therand omP ortio. SOfaProtein donotcontai n Structur a1m o tifsthata rerep eate- with. nt heProtei nsuchasa, e1ixorb p1eatedshe et b1t heem di mensi ona1 fea ture-ofthesePa rtsof. h

eP. Ot inare repea ted f ro mpnemol ec ule toan other Thus, the ter- iS s Om ewhat. famis nomer 42 wh atis theP rim arystruct Ure ofProt eins? WhenaP rote.

ni. COv a lent1 Ymod if ied i tsPrimarys tr uCt UreisCha nged T he primarystructurede.

er. i fst h efinal th feld imen, i ona1 structure oft heProtein Th emodificat iOn diSr. Ptsther old i ng 5 la serine Ha Sasma 11si de chain tha tcanf itina, yreia ti ve1yp0 la renvir0n men t (b. Tr yPtoPh an hasthelar gest si dec hain o f anyofthec ommOnam inOacids, and itten dstore qui re an onpolare nvi ronment (c) Lys ineand arginine.

0t. bas ica11 nolacids e xchanging One fo rthe other wou1dn Otaf fe Ctthesi dechainpKa, ingsignificant, y Simi 1ar reas0n in gaP li eSt0th. Stitu tionOfan OnP ola risole uci neforanonpolarle Ucine 6 G, yci neis freq uent 1yac, nse rvedresidue beca Use its

id. Chain iso Sm allas tofitinSpa Cesth at Wi11, ot accommod atelarger Ones 7 The n1anin eisr ePlacedbyis oleuc ine, therei snot enoug- roomintheena tiveconform, tionforthe lar ge rSidechain 0fth eis

01euc ine c Onsequent ly, ther eisa greatenOu g hchangein, he con forma tionofthe Pr0tei nthat i t0Se SaCtivi ty whengly C ine i Ssubstitutedin tUrnfOri so1eucine, t heP re sencOf t.

eS. a1le rsid e Cha i n1ea dst0 resu. ti0nOft heaCtivec onf0 rmati0n 8 Meatcon sist, 1argelyof animal P rot ei nsa ndfat T hetemP erat. res 9 17. 18 The Conjugate acid baSe pairli 1 lac tasabuffer inthepH ra n ge 1.09 3.91 9 T hey hav eone c harg. atpHextremes, and themo1 ec Ule stendto rep el eachotherly hentheme1. Culatch argei sz ero, the ami noac i dsca nagg regate temo reeasi 1Y 2 0 Theion icdissoc iat

i0. reactio nsofthe f ollowing ami noac ids: aspart i caci d, v. li ne, h ist id ine, serine, and ly si ne 21 The pKa, Or th e i0nizat i0n of th ethi01gr ou pofcys teine is 8 33, So thisami n oaci dC0u1d se rve as a bufferi nthe SH a nd for. SOv ertheP Hr an ge 7 33 9 33 Thea mi n0gro u P S 0, asparagin eandly Sinehave pKav.

1u. Sofs 8 andg 9 5, res pe cti vely; thes ea re a1s0Pos si b1ebuffers, butt heyarebot. near thee nd Of th eirbuff erra nges 22 A tpH 4, the C.

rb. Xylgro UpiSdePro t0 natedtoa carbo xylate, th e sidecha inCarb, Xy 1isst i1 1mor et han 50% Pr oton ated, and bo thamin ogroups are- roton ated At pH 7, b Oththe ca rboxyl gr0Upandthe sidec hainc. rb 0x Y1gr ou paredePr otOnatedto acarbo xy 1ate, and bothami nogroup sare Pro tonated At pH10, both t hecarboxyl grO upandt he sid echai nca rboxy1 groupa redeP ro tonatedtoaC.

rb. Xylate, 0n eof theam inog rouP Si sPr imar, 1ydePro to nat ed, ndtheOt. eram inogroupi sa mi xtureOft he pr oton atedan. depr0tOnate dforms 23 TheP I refers. Otherformi nwhi ch bothcarb oxy 1groU P sarede Proto, atedandb othaminogro.

PS. rOtonate datPH6 9 6 2 3 TheP I refe rstoth

ef. rmin whic h bo thca rboxy1 groUPsared ePrOt on atedand b othaminogr0 UPsPrOton atedatPH, 9 6 24 AtPH1, heC ha rge dgr0 Upsarethe N.

er. i na 1NH3+0nv alinen dt heProtonat edguani di nogrOup Onargi nin e, giV in gzer onetcharg e TheCha rgedgr0UP SatPH7a ret hesaleas PH 1Wi thth ead ditionO fth eCarbOX, 1ategr oupOnt hec te rm ina1 1euC ine, givin g an etchar geof 1 25 Bothp ePtides, PheGl U-er- Met andVa11 lr plcys1eU, haVea Oargeof 1 a tpH1 becauseo ftheProton, tedN-ter mina1aminogro, P A tpH7, the Pe ptideoN the rightha snonet- harge, ecaus eoftheP rOt0nate dN terminal a11 i nogr ouP and he i0n izedC. er min alcarb0 xylate Nega tive charge Thepeptide, nth e1eft has anetcha rg e 0f at, pH7 becauseofthe side chaincar, boxy- ates, oup0 nthe glut amate i naddition to the ch. rge son the N termi- alan, C terminal grou ps 2d (a) Tysine beca.

Se. fth esi deChainam inogroup; (b) ser-nebecause Ofthe1a. kofas ide Chaincarb oXyl; (c) HiStid, ne, becauseof thepresenceofati tra table group int he sidechai n; (d) ASPa rtateb. ca Useofthe Carboxy1 gro UPin thesideC ha in 27 GlyCi neisfrequent1Yu.

ed. Stheba Si So fa bufferinthe aci drange ne ar thepKOf i tscarbOXy1gr 0UP Th euse fu1 bufferrange. Sf omPH 3to3 13 3 4 what is th ePeP t id eBond 28 See Fi gure 3 9 2 The resona nce Structure.

C0. tr ibu te tothePlana r arrangement by8 5

4 6 dnwe Pr ediC tfolding fr0 msequence ?3 5 This 1 eve 1Of sequence

dary structure of proteins 151 Chapter 5

Solubility and Reversibility Globular proteins vary greatly in their solubility in water. Some proteins, such as albumin and globulin, are very soluble, whereas others are insoluble. Generally speaking, globular proteins with a large proportion of hydrophilic groups on the surface of the molecule are soluble in water. Hydrophilic groups include charged groups such as $-NH_3^+$ and $-COO^-$ as well as polar groups such as $-OH$ and $-CONH_2$. Insoluble proteins usually have a large proportion of hydrophobic amino acid residues on the surface of the molecule. Hydrophobic groups include nonpolar groups such as $-CH_3$ and $-C_6H_5$. Insoluble proteins are often found in membranes, where they are anchored in the lipid bilayer by hydrophobic interactions between nonpolar groups on the protein and the hydrocarbon chains of the membrane lipids. Some proteins require the presence of salts or other substances in order to dissolve in water. Others will dissolve in water but will precipitate out of solution if the solution is changed in some way, such as by a change in temperature or pH. This process is usually reversible: The protein will redissolve if the solution is restored to its original conditions. Reversible precipitation is often used in the laboratory to purify proteins.

Denaturation When globular proteins are heated, they undergo a change in conformation called denaturation. The hydrogen bonds and other weak interactions that maintain the native conformation are disrupted by the heat energy, and the protein unfolds into a random coil. Denaturation is usually accompanied by loss of solubility and loss of biological activity. Denaturation can also be caused by changes in pH, which alter the charges on the amino acid residues and disrupt ionic bonds. Denaturation is usually irreversible. Denatured proteins are more susceptible to digestion by proteolytic enzymes, because the unfolded polypeptide chains are more accessible to the enzymes than the compact native molecules.

Specificity Each protein has a unique sequence of amino acids that folds into a unique three-dimensional shape. The specific shape of a protein is responsible for its function. Proteins can bind to other molecules with great specificity because the binding site on the protein has a shape that is complementary to the shape of the molecule that binds to it. Antibodies, for example, are proteins that bind to foreign molecules and help to destroy them. Each antibody binds to a specific foreign molecule, called an antigen. Enzymes are proteins that act as catalysts, speeding up the rate of chemical reactions. Each enzyme binds to a specific substrate and catalyzes a specific reaction. Specificity is the result of the unique shape of the protein molecule.

Classification of Proteins

Proteins are classified into groups on the basis of their shape and solubility. Globular proteins are soluble in water and have a compact, roughly spherical shape. Fibrous proteins are insoluble in water and have a long, thin shape. Globular proteins include most of the proteins with metabolic functions, such as enzymes. Fibrous proteins usually have structural functions, such as collagen in connective tissue and keratin in hair and nails.

Conjugated Proteins

Conjugated proteins are proteins that contain a nonprotein component. The nonprotein component is called a prosthetic group. Conjugated proteins are classified according to the chemical nature of their prosthetic groups. Glycoproteins have carbohydrate prosthetic groups, lipoproteins have lipid prosthetic groups, and metalloproteins have metal ion prosthetic groups. Hemoglobin is a conjugated protein with a heme prosthetic group. Heme contains an iron ion, which binds to oxygen. Hemoglobin is a globular protein that transports oxygen in the blood.

Fibrous Proteins

Fibrous proteins have a long, thin shape. They are insoluble in water. Fibrous proteins usually have structural functions. Collagen is the most abundant protein in mammals. It is found in connective tissue, such as tendons, cartilage, and the organic matrix of bone. Collagen has a triple-helix structure. Keratin is found in hair, nails, feathers, horns, and the outer layer of skin. Keratin has a coiled-coil structure. Fibrin is a protein involved in blood clotting. Actin and myosin are proteins involved in muscle contraction.

Globular Proteins

Globular proteins have a compact, roughly spherical shape. They are usually soluble in water. Globular proteins usually have metabolic functions. Enzymes are globular proteins that act as catalysts. Hemoglobin is a globular protein that transports oxygen in the blood. Antibodies are globular proteins that bind to foreign molecules and help to destroy them. Some hormones are globular proteins. Globular proteins are more diverse in structure and function than fibrous proteins.

Protein Structure

The structure of a protein is described at four levels: primary, secondary, tertiary, and quaternary. The primary structure is the sequence of amino acids in the polypeptide chain. The secondary structure is the local folding of the polypeptide chain into regular patterns, such as the α-helix and the β-pleated sheet. The tertiary structure is the overall three-dimensional shape of the polypeptide chain. The quaternary structure is the arrangement of two or more polypeptide chains in a protein that has more than one chain.

Primary Structure

The primary structure of a protein is the sequence of amino acids in the polypeptide chain. The amino acids are joined by peptide bonds. The primary structure is determined by the genetic code. The primary structure is the most important level of protein structure, because it determines the higher levels of structure. A change in the primary structure can affect the function of the protein. For example, sickle cell anemia is caused by a single amino acid substitution in the hemoglobin molecule.

Secondary Structure

The secondary structure of a protein is the local folding of the polypeptide chain into regular patterns. The most common secondary structures are the α-helix and the β-pleated sheet. The α-helix is a coiled structure held together by hydrogen bonds between the amino acids. The β-pleated sheet is a sheet-like structure held together by hydrogen bonds between adjacent polypeptide chains. Some proteins have regions of α-helix and β-pleated sheet, while others have neither.

Tertiary Structure

The tertiary structure of a protein is the overall three-dimensional shape of the polypeptide chain. The tertiary structure is determined by interactions between the amino acid side chains. These interactions include hydrogen bonds, ionic bonds, hydrophobic interactions, and disulfide bridges. The tertiary structure is the most complex level of protein structure. It is the level that is most affected by changes in the environment, such as temperature and pH.

Quaternary Structure

The quaternary structure of a protein is the arrangement of two or more polypeptide chains in a protein that has more than one chain. The individual polypeptide chains are called subunits. The quaternary structure is held together by the same types of interactions that hold the tertiary structure together. Hemoglobin is an example of a protein with quaternary structure. It has four subunits: two α-chains and two β-chains.